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(54) Title: HUMAN GENES AND GENE EXPRESSION PRODUCTS ISOLATED FROM HUMAN PROSTATE

(57) Abstract: This invention relates to novel human polynucleotides and variants thereof, their encoded polypeptides and variants thereof, to genes corresponding to these polynucleotides and to proteins expressed by the genes. The invention also relates to diagnostics and therapeutics comprising such novel human polynucleotides, their corresponding genes or gene products, including probes, antisense nucleotides, and antibodies. The polynucleotides of the invention correspond to a polynucleotide comprising the sequence information of at least one of SEQ ID NOS:1-1485. The polypeptides of the invention correspond to a polypeptide comprising the amino acid sequence information of at least one of SEQ ID NOS:1486-1542.

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# **HUMAN GENES AND GENE EXPRESSION PRODUCTS ISOLATED FROM HUMAN PROSTATE**

## Field of the Invention

5       The present invention relates to polynucleotides of human origin, particularly in human prostate, and the encoded gene products.

## Background of the Invention

Identification of novel polynucleotides, particularly those that encode an expressed gene product, is important in the advancement of drug discovery, diagnostic technologies, and the  
10       understanding of the progression and nature of complex diseases such as cancer. Identification of genes expressed in different cell types isolated from sources that differ in disease state or stage, developmental stage, exposure to various environmental factors, the tissue of origin, the species from which the tissue was isolated, and the like is key to identifying the genetic factors that are responsible for the phenotypes associated with these various differences.

15       This invention provides novel human polynucleotides, the polypeptides encoded by these polynucleotides, and the genes and proteins corresponding to these novel polynucleotides.

## Summary of the Invention

This invention relates to novel human polynucleotides and variants thereof, their encoded polypeptides and variants thereof, to genes corresponding to these polynucleotides and to proteins  
20       expressed by the genes. The invention also relates to diagnostics and therapeutics comprising such novel human polynucleotides, their corresponding genes or gene products, including probes, antisense nucleotides, and antibodies. The polynucleotides of the invention correspond to a polynucleotide comprising the sequence information of at least one of SEQ ID NOS:1-1485. The polypeptides of the invention correspond to a polypeptide comprising the amino acid sequence information of at least one  
25       of SEQ ID NOS:1486-1542.

Accordingly, in one aspect, the invention provides an isolated polynucleotide comprising a nucleotide sequence which hybridizes under stringent conditions to a sequence selected from the group consisting of SEQ ID NOS: 1-1485.

30       In another aspect, the invention provides an isolated polynucleotide comprising at least 15 contiguous nucleotides of a nucleotide sequence having at least 90% sequence identity to a sequence selected from the group consisting of SEQ ID NOS:1-1485, a degenerate variant of SEQ ID NOS:1-1485, an antisense of SEQ ID NOS:1-1485, and a complement of SEQ ID NOS:1-1485.

In another aspect, the invention provides an isolated polynucleotide comprising at least 15 contiguous nucleotides of a nucleotide sequence selected from the group consisting of SEQ ID  
35       NOS:1-1485, a degenerate variant of SEQ ID NOS:1-1485, an antisense of SEQ ID NOS:1-1485, and a complement of SEQ ID NOS:1-1485. In specific embodiments, the polynucleotide comprises at

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least 100 contiguous nucleotides of the nucleotide sequence. In other specific embodiments, the polynucleotide comprises at least 200 contiguous nucleotides of the nucleotide sequence.

In another aspect, the invention provides An isolated polynucleotide comprising a nucleotide sequence of at least 90% sequence identity to a sequence selected from the group consisting of SEQ ID NOS: 1-1485, a degenerate variant of SEQ ID NOS: 1-1485, an antisense of SEQ ID NOS: 1-1485, and a complement of SEQ ID NOS: 1-1485. In specific embodiments, the polynucleotide comprises a nucleotide sequence of at least 95% sequence identity to the selected nucleotide sequence. In other specific embodiments, the polynucleotide comprises a nucleotide sequence that is identical to the selected nucleotide sequence.

In another aspect, the invention provides a polynucleotide comprising a nucleotide sequence of an insert contained in a clone deposited as NRRL Accession No. B-30523, B-30524, B-30525, B-30526, B-30527, B-30528, B-30529, or B-30581.

In another aspect, the invention provides an isolated cDNA obtained by the process of amplification using a polynucleotide comprising at least 15 contiguous nucleotides of a nucleotide sequence selected from the group consisting of SEQ ID NOS: 1-1485. In specific embodiments, the polynucleotide comprises at least 25 contiguous nucleotides of the selected nucleotide sequence. In other specific embodiments, the polynucleotide comprises at least 100 contiguous nucleotides of the selected nucleotide sequence. In some embodiments, the amplification is by polymerase chain reaction (PCR) amplification.

In another aspect, the invention provides an isolated recombinant host cell containing a polynucleotide of the invention.

In another aspect, the invention provides an isolated vector comprising a polynucleotide of the invention.

In another aspect, the invention provides a method for producing a polypeptide, the method comprising the steps of culturing a recombinant host cell containing a polynucleotide of the invention under conditions suitable for the expression of an encoded polypeptide and recovering the polypeptide from the host cell culture.

In another aspect, the invention provides an isolated polypeptide encoded by a polynucleotide of the invention.

In another aspect, the invention provides an isolated polypeptide comprising an amino acid sequence selected from the group consisting of SEQ ID NOS: 1486-1542.

In another aspect, the invention provides an antibody that specifically binds a polypeptide of the invention.

In another aspect, the invention provides a method of detecting differentially expressed genes correlated with a cancerous state of a mammalian cell, the method comprising the step of detecting at least one differentially expressed gene product in a test sample derived from a cell suspected of being

cancerous, where the gene product is encoded by a gene comprising an identifying sequence of at least one of SEQ ID NOS:1-1485. Detection of the differentially expressed gene product is correlated with a cancerous state of the cell from which the test sample was derived.

In another aspect, the invention provides a method of detecting differentially expressed genes correlated with a cancerous state of a mammalian cell, the method comprising the step of detecting at least one differentially expressed gene product in a test sample derived from a cell suspected of being cancerous, where the gene product comprises an amino acid sequence selected from the group consisting of SEQ ID NOS:1486-1542. Detection of the differentially expressed gene product is correlated with a cancerous state of the cell from which the test sample was derived.

In another aspect, the invention provides a library of polynucleotides, wherein at least one of the polynucleotides comprises the sequence information of a polynucleotide of the invention. In specific embodiments, the library is provided on a nucleic acid array. In some embodiments, the library is provided in a computer-readable format.

In another aspect, the invention provides a method of inhibiting tumor growth by modulating expression of a gene product, the gene product being encoded by a gene identified by a sequence selected from the group consisting of SEQ ID NOS:1-1485.

In another aspect, the invention provides a method of inhibiting tumor growth by modulating expression of a gene product, the gene product comprising an amino acid sequence selected from the group consisting of SEQ ID NOS: 1486-1542.

These and other objects, advantages, and features of the invention will become apparent to those persons skilled in the art upon reading the details of the invention as more fully described below.

#### Detailed Description of the Invention

Before the present invention is described, it is to be understood that this invention is not limited to particular embodiments described, as such may, of course, vary. It is also to be understood that the terminology used herein is for the purpose of describing particular embodiments only, and is not intended to be limiting.

Unless defined otherwise, all technical and scientific terms used herein have the same meaning as commonly understood by one of ordinary skill in the art to which this invention belongs. Although any methods and materials similar or equivalent to those described herein can be used in the practice or testing of the present invention, the preferred methods and materials are now described.

All publications and patent applications cited in this specification are herein incorporated by reference as if each individual publication or patent application were specifically and individually indicated to be incorporated by reference. The citation of any publication is for its disclosure prior to the filing date and should not be construed as an admission that the present invention is not entitled to antedate such publication by virtue of prior invention.



It must be noted that as used herein and in the appended claims, the singular forms "a," "and," and "the" include plural referents unless the context clearly dictates otherwise. Thus, for example, reference to "a polynucleotide" includes a plurality of such polynucleotides and reference to "the colon cancer cell" includes reference to one or more cells and equivalents thereof known to those skilled in the art, and so forth.

The publications and applications discussed herein are provided solely for their disclosure prior to the filing date of the present application. Nothing herein is to be construed as an admission that the present invention is not entitled to antedate such publication by virtue of prior invention. Further, the dates of publication provided may be different from the actual publication dates which may need to be independently confirmed.

### Definitions

The terms "polynucleotide" and "nucleic acid," used interchangeably herein, refer to a polymeric forms of nucleotides of any length, either ribonucleotides or deoxynucleotides. Thus, these terms include, but are not limited to, single-, double-, or multi-stranded DNA or RNA, genomic DNA, cDNA, DNA-RNA hybrids, branched nucleic acid (see, e.g., U.S. Pat. Nos. 5,124,246; 5,710,264; and 5,849,481), or a polymer comprising purine and pyrimidine bases or other natural, chemically or biochemically modified, non-natural, or derivatized nucleotide bases. These terms further include, but are not limited to, mRNA or cDNA that comprise intronic sequences (see, e.g., Niwa et al. (1999) Cell 99(7):691-702). The backbone of the polynucleotide can comprise sugars and phosphate groups (as may typically be found in RNA or DNA), or modified or substituted sugar or phosphate groups. Alternatively, the backbone of the polynucleotide can comprise a polymer of synthetic subunits such as phosphoramidites and thus can be an oligodeoxynucleoside phosphoramidate or a mixed phosphoramidate-phosphodiester oligomer. Peyrottes et al. (1996) Nucl. Acids Res. 24:1841-1848; Chaturvedi et al. (1996) Nucl. Acids Res. 24:2318-2323. A polynucleotide may comprise modified nucleotides, such as methylated nucleotides and nucleotide analogs, uracyl, other sugars, and linking groups such as fluororibose and thioate, and nucleotide branches. The sequence of nucleotides may be interrupted by non-nucleotide components. A polynucleotide may be further modified after polymerization, such as by conjugation with a labeling component. Other types of modifications included in this definition are caps, substitution of one or more of the naturally occurring nucleotides with an analog, and introduction of means for attaching the polynucleotide to proteins, metal ions, labeling components, other polynucleotides, or a solid support.

The terms "polypeptide" and "protein," used interchangeably herein, refer to a polymeric form of amino acids of any length, which can include coded and non-coded amino acids, chemically or biochemically modified or derivatized amino acids, and polypeptides having modified peptide backbones. The term includes fusion proteins, including, but not limited to, fusion proteins with a

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heterologous amino acid sequence, fusions with heterologous and homologous leader sequences, with or without N-terminal methionine residues; immunologically tagged proteins; and the like.

"Diagnosis" as used herein generally includes determination of a subject's susceptibility to a disease or disorder, determination as to whether a subject is presently affected by a disease or disorder, prognosis of a subject affected by a disease or disorder (e.g., identification of pre-metastatic or metastatic cancerous states, stages of cancer, or responsiveness of cancer to therapy), and therapeutics (e.g., monitoring a subject's condition to provide information as to the effect or efficacy of therapy).

"Sample" or "biological sample" as used herein encompasses a variety of sample types, and are generally meant to refer to samples of biological fluids or tissues, particularly samples obtained from tissues, especially from cells of the type associated with a disease or condition for which a diagnostic application is designed (e.g., ductal adenocarcinoma), and the like. "Sample" or "biological sample" are meant to encompass blood and other liquid samples of biological origin, solid tissue samples, such as a biopsy specimen or tissue cultures or cells derived therefrom and the progeny thereof. These terms encompass samples that have been manipulated in any way after their procurement as well as derivatives and fractions of samples, where the samples may be manipulated by, for example, treatment with reagents, solubilization, or enrichment for certain components. The terms also encompass clinical samples, and also includes cells in cell culture, cell supernatants, cell lysates, serum, plasma, biological fluids, and tissue samples. Where the sample is solid tissue, the cells of the tissue can be dissociated or tissue sections can be analyzed.

The terms "treatment," "treating," "treat" and the like are used herein to generally refer to obtaining a desired pharmacologic and/or physiologic effect. The effect may be prophylactic in terms of completely or partially preventing a disease or symptom thereof and/or may be therapeutic in terms of a partial or complete stabilization or cure for a disease and/or adverse effect attributable to the disease. "Treatment" as used herein covers any treatment of a disease in a mammal, particularly a human, and includes: (a) preventing the disease or symptom from occurring in a subject which may be predisposed to the disease or symptom but has not yet been diagnosed as having it; (b) inhibiting the disease symptom, i.e., arresting its development; or relieving the disease symptom, i.e., causing regression of the disease or symptom.

The terms "individual," "subject," "host," and "patient," used interchangeably herein and refer to any mammalian subject for whom diagnosis, treatment, or therapy is desired, particularly humans. Other subjects may include cattle, dogs, cats, guinea pigs, rabbits, rats, mice, horses, and so on.

As used herein the term "isolated" refers to a polynucleotide, a polypeptide, an antibody, or a host cell that is in an environment different from that in which the polynucleotide, the polypeptide, the antibody, or the host cell naturally occurs. A polynucleotide, a polypeptide, an antibody, or a host cell which is isolated is generally substantially purified. As used herein, the term "substantially purified" refers to a compound (e.g., either a polynucleotide or a polypeptide or an antibody) that is removed

from its natural environment and is at least 60% free, preferably 75% free, and most preferably 90% free from other components with which it is naturally associated. Thus, for example, a composition containing A is "substantially free of" B when at least 85% by weight of the total A+B in the composition is A. Preferably, A comprises at least about 90% by weight of the total of A+B in the composition, more preferably at least about 95% or even 99% by weight.

A "host cell," as used herein, refers to a microorganism or a eukaryotic cell or cell line cultured as a unicellular entity which can be, or has been, used as a recipient for a recombinant vector or other transfer polynucleotides, and include the progeny of the original cell which has been transfected. It is understood that the progeny of a single cell may not necessarily be completely identical in morphology or in genomic or total DNA complement as the original parent, due to natural, accidental, or deliberate mutation.

The terms "cancer," "neoplasm," "tumor," and "carcinoma," are used interchangeably herein to refer to cells which exhibit relatively autonomous growth, so that they exhibit an aberrant growth phenotype characterized by a significant loss of control of cell proliferation. In general, cells of interest for detection or treatment in the present application include precancerous (e.g., benign), malignant, metastatic, and non-metastatic cells. Detection of cancerous cell is of particular interest.

The use of "e", as in 10e-3, indicates that the number to the left of "e" is raised to the power of the number to the right of "e" (thus, 10e-3 is  $10^{-3}$ ).

The term "heterologous" as used herein in the context of, for example, heterologous nucleic acid or amino acid sequences, heterologous polypeptides, or heterologous nucleic acid, is meant to refer to material that originates from a source different from that with which it is joined or associated. For example, two DNA sequences are heterologous to one another if the sequences are from different genes or from different species. A recombinant host cell containing a sequence that is heterologous to the host cell can be, for example, a bacterial cell containing a sequence encoding a human polypeptide.

The invention relates to polynucleotides comprising the disclosed nucleotide sequences, to full length cDNA, mRNA, genomic sequences, and genes corresponding to these sequences and degenerate variants thereof, and to polypeptides encoded by the polynucleotides of the invention and polypeptide variants. The following detailed description describes the polynucleotide compositions encompassed by the invention, methods for obtaining cDNA or genomic DNA encoding a full-length gene product, expression of these polynucleotides and genes, identification of structural motifs of the polynucleotides and genes, identification of the function of a gene product encoded by a gene corresponding to a polynucleotide of the invention, use of the provided polynucleotides as probes and in mapping and in tissue profiling, use of the corresponding polypeptides and other gene products to raise antibodies, and use of the polynucleotides and their encoded gene products for therapeutic and diagnostic purposes.

# Polynucleotide Compositions

The present invention provides isolated polynucleotides that represent genes that are differentially expressed in human cancer cells. The polynucleotides, as well as polypeptides encoded thereby, find use in a variety of therapeutic and diagnostic methods.

5       The scope of the invention with respect to compositions containing the isolated polynucleotides useful in the methods described herein includes, but is not necessarily limited to, polynucleotides having a sequence set forth in any one of the polynucleotide sequences provided herein; polynucleotides obtained from the biological materials described herein or other biological sources (particularly human sources) by hybridization under stringent conditions (particularly  
10       conditions of high stringency); genes corresponding to the provided polynucleotides; cDNAs corresponding to the provided polynucleotides; variants of the provided polynucleotides and their corresponding genes, particularly those variants that retain a biological activity of the encoded gene product (e.g., a biological activity ascribed to a gene product corresponding to the provided polynucleotides as a result of the assignment of the gene product to a protein family(ies) and/or  
15       identification of a functional domain present in the gene product). Other nucleic acid compositions contemplated by and within the scope of the present invention will be readily apparent to one of ordinary skill in the art when provided with the disclosure here. "Polynucleotide" and "nucleic acid" as used herein with reference to nucleic acids of the composition is not intended to be limiting as to the length or structure of the nucleic acid unless specifically indicated.

20       The invention features polynucleotides that represent genes that are expressed in human tissue, specifically human breast tissue, particularly polynucleotides that are differentially expressed in cancerous breast cells. Nucleic acid compositions described herein of particular interest are at least about 15 bp in length, at least about 30 bp in length, at least about 50 bp in length, at least about 100 bp, at least about 200 bp in length, at least about 300 bp in length, at least about 500 bp in length, at  
25       least about 800 bp in length, at least about 1 kb in length, at least about 2.0 kb in length, at least about 3.0 kb in length, at least about 5 kb in length, at least about 10 kb in length, at least about 50kb in length and are usually less than about 200 kb in length. These polynucleotides (or polynucleotide fragments) have uses that include, but are not limited to, diagnostic probes and primers as starting materials for probes and primers, as discussed herein.

30       The subject polynucleotides usually comprise a sequence set forth in any one of the polynucleotide sequences provided herein, for example, in the sequence listing, incorporated by reference in a table (e.g. by an NCBI accession number), a cDNA deposited at the A.T.C.C., or a fragment or variant thereof. A "fragment" or "portion" of a polynucleotide is a contiguous sequence of residues at least about 10 nt to about 12 nt, 15 nt, 16 nt, 18 nt or 20 nt in length, usually at least  
35       about 22 nt, 24 nt, 25 nt, 30 nt, 40 nt, 50 nt, 60nt, 70 nt, 80 nt, 90 nt, 100 nt to at least about 150 nt, 200 nt, 250 nt, 300 nt, 350 nt, 400 nt, 500 nt, 800 nt or up to about 1000 nt, 1500 or 2000 nt in

length. In some embodiments, a fragment of a polynucleotide is the coding sequence of a polynucleotide. A fragment of a polynucleotide may start at position 1 (i.e. the first nucleotide) of a nucleotide sequence provided herein, or may start at about position 10, 20, 30, 50, 75, 100, 150, 200, 250, 300, 350, 400, 450, 500, 600, 700, 800, 900, 1000, 1500 or 2000, or an ATG translational initiation codon of a nucleotide sequence provided herein. In this context "about" includes the particularly recited value or a value larger or smaller by several (5, 4, 3, 2, or 1) nucleotides. The described polynucleotides and fragments thereof find use as hybridization probes, PCR primers, BLAST probes, or as an identifying sequence, for example.

The subject nucleic acids may be variants or degenerate variants of a sequence provided herein. In general, a variants of a polynucleotide provided herein have a fragment of sequence identity that is greater than at least about 65%, greater than at least about 70%, greater than at least about 75%, greater than at least about 80%, greater than at least about 85%, or greater than at least about 90%, 95%, 96%, 97%, 98%, 99% or more (i.e. 100%) as compared to an identically sized fragment of a provided sequence. as determined by the Smith-Waterman homology search algorithm as implemented in MPSRCH program (Oxford Molecular). For the purposes of this invention, a preferred method of calculating percent identity is the Smith-Waterman algorithm. Global DNA sequence identity should be greater than 65% as determined by the Smith-Waterman homology search algorithm as implemented in MPSRCH program (Oxford Molecular) using an gap search with the following search parameters: gap open penalty, 12; and gap extension penalty, 1.

The subject nucleic acid compositions include full-length cDNAs or mRNAs that encompass an identifying sequence of contiguous nucleotides from any one of the polynucleotide sequences provided herein.

As discussed above, the polynucleotides useful in the methods described herein also include polynucleotide variants having sequence similarity or sequence identity. Nucleic acids having sequence similarity are detected by hybridization under low stringency conditions, for example, at 50°C and 10XSSC (0.9 M saline/0.09 M sodium citrate) and remain bound when subjected to washing at 55°C in 1XSSC. Sequence identity can be determined by hybridization under high stringency conditions, for example, at 50°C or higher and 0.1XSSC (9 mM saline/0.9 mM sodium citrate). Hybridization methods and conditions are well known in the art, see, e.g., USPN 5,707,829. Nucleic acids that are substantially identical to the provided polynucleotide sequences, e.g. allelic variants, genetically altered versions of the gene, etc., bind to the provided polynucleotide sequences under stringent hybridization conditions. By using probes, particularly labeled probes of DNA sequences, one can isolate homologous or related genes. The source of homologous genes can be any species, e.g. primate species, particularly human; rodents, such as rats and mice; canines, felines, bovines, ovines, equines, yeast, nematodes, etc.

In one embodiment, hybridization is performed using a fragment of at least 15 contiguous nucleotides (nt) of at least one of the polynucleotide sequences provided herein. That is, when at least 15 contiguous nt of one of the disclosed polynucleotide sequences is used as a probe, the probe will preferentially hybridize with a nucleic acid comprising the complementary sequence, allowing the identification and retrieval of the nucleic acids that uniquely hybridize to the selected probe. Probes from more than one polynucleotide sequence provided herein can hybridize with the same nucleic acid if the cDNA from which they were derived corresponds to one mRNA.

Polynucleotides contemplated for use in the invention also include those having a sequence of naturally occurring variants of the nucleotide sequences (e.g., degenerate variants (e.g., sequences that encode the same polypeptides but, due to the degenerate nature of the genetic code, different in nucleotide sequence), allelic variants, etc.). Variants of the polynucleotides contemplated by the invention are identified by hybridization of putative variants with nucleotide sequences disclosed herein, preferably by hybridization under stringent conditions. For example, by using appropriate wash conditions, variants of the polynucleotides described herein can be identified where the allelic variant exhibits at most about 25-30% base pair (bp) mismatches relative to the selected polynucleotide probe. In general, allelic variants contain 15-25% bp mismatches, and can contain as little as even 5-15%, or 2-5%, or 1-2% bp mismatches, as well as a single bp mismatch.

The invention also encompasses homologs corresponding to any one of the polynucleotide sequences provided herein, where the source of homologous genes can be any mammalian species, e.g., primate species, particularly human; rodents, such as rats; canines, felines, bovines, ovines, equines, yeast, nematodes, etc. Between mammalian species, e.g., human and mouse, homologs generally have substantial sequence similarity, e.g., at least 75% sequence identity, usually at least 80%, at least 85, at least 90%, at least 95%, at least 96%, at least 97%, at least 98%, at least 99% or even 100% identity between nucleotide sequences. Sequence similarity is calculated based on a reference sequence, which may be a subset of a larger sequence, such as a conserved motif, coding region, flanking region, etc. A reference sequence will usually be at least about a fragment of a polynucleotide sequence and may extend to the complete sequence that is being compared. Algorithms for sequence analysis are known in the art, such as gapped BLAST, described in Altschul, et al. *Nucleic Acids Res.* (1997) 25:3389-3402, or TeraBLAST available from TimeLogic Corp. (Crystal Bay, Nevada).

Moreover, representative examples of polynucleotide fragments of the invention (useful, for example, as probes), include, for example, fragments comprising, or alternatively consisting of, a sequence from about nucleotide number 1-50, 51-100, 101-150, 151-200, 201-250, 251-300, 301-350, 351-400, 401-450, 451-500, 501-550, 551-600, 601-700, 701-750, 751-800, 800-850, 851-900, 901-950, 951-1000, 1001-1050, 1051-1100, 1101-1150, 1151-1200, 1201-1250, 1251-1300, 1301-1350, 1351-1400, 1401-1450, 1451-1500, 1501-1550, 1551-1600, 1601-1650, 1651-1700, 1701-1750,

1751-1800, 1801-1850, 1851-1900, 1901-1950, 1951-2000, 2001-2050, 2051-2100, 2101-2150, 2151-2200, 2201-2250, 2251-2300, 2301-2350, 2351-2400, 2401-2450, 2451-2500, 2501-2550, 2551-2600, 2601-2650, 2651-2700, 2701-2750, 2751-2800, 2801-2850, 2851-2900, 2901-2950, 2951-3000, 3001-3050, 3051-3100, 3101-3150, 3151-3200, 3201-3250, 3251-3300, 3301-3350, 3351-3400, 3401-3450, 3451-3500, 3501-3550, 3551-3600, 3601-3650, 3651-3700, 3701-3750, 3751-3800, 3801-3850, 3851-3900, 3901-3950, 3951-4000, 4001-4050, 4051-4100, 4101-4150, 4151-4200, 4201-4250, 4251-4300, 4301-4350, 4351-4400, 4401-4450, 4451-4500, 4501-4550, 4551-4600, 4601-4650, 4651-4700, 4701-4750, 4751-4800, 4801-4850, 4851-4900, 4901-4950, 4951-5000, 5001-5050, 5051-5100, 5101-5150, 5151-5200, 5201-5250, 5251-5300, 5301-5350, 5351-5400, 5401-5450, 5451-5500, 5501-5550, 5551-5600, 5601-5650, 5651-5700, 5701-5750, 5751-5800, 5801-5850, 5851-5900, 5901-5950, 5951-6000, 6001-6050, 6051-6100, 6101-6150, and 6151 of a subject nucleic acid, or the complementary strand thereto. In this context "about" includes the particularly recited range or a range larger or smaller by several (5, 4, 3, 2, or 1) nucleotides, at either terminus or at both termini. In some embodiments, these fragments encode a polypeptide which has a functional activity (e.g., biological activity) whereas in other embodiments, these fragments are probes, or starting materials for probes. Polynucleotides which hybridize to one or more of these nucleic acid molecules under stringent hybridization conditions or alternatively, under lower stringency conditions, are also encompassed by the invention, as are polypeptides encoded by these polynucleotides or fragments.

The subject nucleic acids can be cDNAs or genomic DNAs, as well as fragments thereof, particularly fragments that encode a biologically active gene product and/or are useful in the methods disclosed herein (e.g., in diagnosis, as a unique identifier of a differentially expressed gene of interest, etc.). The term "cDNA" as used herein is intended to include all nucleic acids that share the arrangement of sequence elements found in native mature mRNA species, where sequence elements are exons and 3' and 5' non-coding regions. Normally mRNA species have contiguous exons, with the intervening introns, when present, being removed by nuclear RNA splicing, to create a continuous open reading frame encoding a polypeptide. mRNA species can also exist with both exons and introns, where the introns may be removed by alternative splicing. Furthermore it should be noted that different species of mRNAs encoded by the same genomic sequence can exist at varying levels in a cell, and detection of these various levels of mRNA species can be indicative of differential expression of the encoded gene product in the cell.

A genomic sequence of interest comprises the nucleic acid present between the initiation codon and the stop codon, as defined in the listed sequences, including all of the introns that are normally present in a native chromosome. It can further include the 3' and 5' untranslated regions found in the mature mRNA. It can further include specific transcriptional and translational regulatory sequences, such as promoters, enhancers, etc., including about 1 kb, but possibly more, of flanking

genomic DNA at either the 5' and 3' end of the transcribed region. The genomic DNA can be isolated as a fragment of 100 kbp or smaller, and substantially free of flanking chromosomal sequence. The genomic DNA flanking the coding region, either 3' and 5', or internal regulatory sequences as sometimes found in introns, contains sequences required for proper tissue, stage-specific, or disease-state specific expression.

The nucleic acid compositions of the subject invention can encode all or a part of the naturally-occurring polypeptides. Double or single stranded fragments can be obtained from the DNA sequence by chemically synthesizing oligonucleotides in accordance with conventional methods, by restriction enzyme digestion, by PCR amplification, *etc.*

Probes specific to the polynucleotides described herein can be generated using the polynucleotide sequences disclosed herein. The probes are usually a fragment of a polynucleotide sequences provided herein. The probes can be synthesized chemically or can be generated from longer polynucleotides using restriction enzymes. The probes can be labeled, for example, with a radioactive, biotinylated, or fluorescent tag. Preferably, probes are designed based upon an identifying sequence of any one of the polynucleotide sequences provided herein. More preferably, probes are designed based on a contiguous sequence of one of the subject polynucleotides that remain unmasked following application of a masking program for masking low complexity (*e.g.*, XBLAST, RepeatMasker, *etc.*) to the sequence., *i.e.*, one would select an unmasked region, as indicated by the polynucleotides outside the poly-n stretches of the masked sequence produced by the masking program.

The polynucleotides of interest in the subject invention are isolated and obtained in substantial purity, generally as other than an intact chromosome. Usually, the polynucleotides, either as DNA or RNA, will be obtained substantially free of other naturally-occurring nucleic acid sequences that they are usually associated with, generally being at least about 50%, usually at least about 90% pure and are typically "recombinant", *e.g.*, flanked by one or more nucleotides with which it is not normally associated on a naturally occurring chromosome.

The polynucleotides described herein can be provided as a linear molecule or within a circular molecule, and can be provided within autonomously replicating molecules (vectors) or within molecules without replication sequences. Expression of the polynucleotides can be regulated by their own or by other regulatory sequences known in the art. The polynucleotides can be introduced into suitable host cells using a variety of techniques available in the art, such as transferrin polycation-mediated DNA transfer, transfection with naked or encapsulated nucleic acids, liposome-mediated DNA transfer, intracellular transportation of DNA-coated latex beads, protoplast fusion, viral infection, electroporation, gene gun, calcium phosphate-mediated transfection, and the like.

The nucleic acid compositions described herein can be used to, for example, produce polypeptides, as probes for the detection of mRNA in biological samples (*e.g.*, extracts of human



cells) or cDNA produced from such samples, to generate additional copies of the polynucleotides, to generate ribozymes or antisense oligonucleotides, and as single stranded DNA probes or as triple-strand forming oligonucleotides. The probes described herein can be used to, for example, determine the presence or absence of any one of the polynucleotide provided herein or variants thereof in a sample. These and other uses are described in more detail below. The subject nucleic acid compositions can be used, for example, to produce polypeptides, as probes for the detection of mRNA of the invention in biological samples (e.g., extracts of human cells) to generate additional copies of the polynucleotides, to generate ribozymes or antisense oligonucleotides, and as single stranded DNA probes or as triple-strand forming oligonucleotides. The probes described herein can be used to, for example, determine the presence or absence of the polynucleotide sequences as shown in SEQ ID NOS:1-1485 or variants thereof in a sample. These and other uses are described in more detail below.

#### Use of Polynucleotides to Obtain Full-Length cDNA, Gene, and Promoter Region

In one embodiment, the polynucleotides are useful as starting materials to construct larger molecules. In one example, the polynucleotides of the invention are used to construct polynucleotides that encode a larger polypeptide (e.g., up to the full-length native polypeptide as well as fusion proteins comprising all or a portion of the native polypeptide) or may be used to produce haptens of the polypeptide (e.g., polypeptides useful to generate antibodies).

In one particular example, the polynucleotides of the invention are used to make or isolate cDNA molecules encoding all or portion of a naturally-occurring polypeptide. Full-length cDNA molecules comprising the disclosed polynucleotides are obtained as follows. A polynucleotide having a sequence of one of SEQ ID NOS:1-1485, or a portion thereof comprising at least 12, 15, 18, or 20 nt, is used as a hybridization probe to detect hybridizing members of a cDNA library using probe design methods, cloning methods, and clone selection techniques such as those described in USPN 5,654,173. Libraries of cDNA are made from selected tissues, such as normal or tumor tissue, or from tissues of a mammal treated with, for example, a pharmaceutical agent. Preferably, the tissue is the same as the tissue from which the polynucleotides of the invention were isolated, as both the polynucleotides described herein and the cDNA represent expressed genes. Most preferably, the cDNA library is made from the biological material described herein in the Examples. The choice of cell type for library construction can be made after the identity of the protein encoded by the gene corresponding to the polynucleotide of the invention is known. This will indicate which tissue and cell types are likely to express the related gene, and thus represent a suitable source for the mRNA for generating the cDNA. Where the provided polynucleotides are isolated from cDNA libraries, the libraries are prepared from mRNA of human prostate cells, more preferably, human prostate cancer cells

Techniques for producing and probing nucleic acid sequence libraries are described, for example, in Sambrook et al., *Molecular Cloning: A Laboratory Manual*, 2nd Ed., (1989) Cold Spring

Harbor Press, Cold Spring Harbor, NY. The cDNA can be prepared by using primers based on polynucleotides comprising a sequence of SEQ ID NOS:1-1485. In one embodiment, the cDNA library can be made from only poly-adenylated mRNA. Thus, poly-T primers can be used to prepare cDNA from the mRNA.

5 Members of the library that are larger than the provided polynucleotides, and preferably that encompass the complete coding sequence of the native message, are obtained. In order to confirm that the entire cDNA has been obtained, RNA protection experiments are performed as follows. Hybridization of a full-length cDNA to an mRNA will protect the RNA from RNase degradation. If the cDNA is not full length, then the portions of the mRNA that are not hybridized will be subject to  
10 RNase degradation. This is assayed, as is known in the art, by changes in electrophoretic mobility on polyacrylamide gels, or by detection of released monoribonucleotides. Sambrook et al., Molecular Cloning: A Laboratory Manual, 2nd Ed., (1989) Cold Spring Harbor Press, Cold Spring Harbor, NY. In order to obtain additional sequences 5' to the end of a partial cDNA, 5' RACE (PCR Protocols: A Guide to Methods and Applications, (1990) Academic Press, Inc.) can be performed.

15 Genomic DNA is isolated using the provided polynucleotides in a manner similar to the isolation of full-length cDNAs. Briefly, the provided polynucleotides, or portions thereof, are used as probes to libraries of genomic DNA. Preferably, the library is obtained from the cell type that was used to generate the polynucleotides of the invention, but this is not essential. Most preferably, the genomic DNA is obtained from the biological material described herein in the Examples. Such  
20 libraries can be in vectors suitable for carrying large segments of a genome, such as P1 or YAC, as described in detail in Sambrook et al., supra, 9.4-9.30. In addition, genomic sequences can be isolated from human BAC libraries, which are commercially available from Research Genetics, Inc., Huntsville, Alabama, USA, for example. In order to obtain additional 5' or 3' sequences, chromosome walking is performed, as described in Sambrook et al., such that adjacent and overlapping fragments  
25 of genomic DNA are isolated. These are mapped and pieced together, as is known in the art, using restriction digestion enzymes and DNA ligase.

Using the polynucleotide sequences of the invention, corresponding full-length genes can be isolated using both classical and PCR methods to construct and probe cDNA libraries. Using either method, Northern blots, preferably, are performed on a number of cell types to determine which cell  
30 lines express the gene of interest at the highest level. Classical methods of constructing cDNA libraries are taught in Sambrook et al., supra. With these methods, cDNA can be produced from mRNA and inserted into viral or expression vectors. Typically, libraries of mRNA comprising poly(A) tails can be produced with poly(T) primers. Similarly, cDNA libraries can be produced using the instant sequences as primers.

35 PCR methods are used to amplify the members of a cDNA library that comprise the desired insert. In this case, the desired insert will contain sequence from the full length cDNA that

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corresponds to the instant polynucleotides. Such PCR methods include gene trapping and RACE methods. Gene trapping entails inserting a member of a cDNA library into a vector. The vector then is denatured to produce single stranded molecules. Next, a substrate-bound probe, such as a biotinylated oligo, is used to trap cDNA inserts of interest. Biotinylated probes can be linked to an avidin-bound solid substrate. PCR methods can be used to amplify the trapped cDNA. To trap sequences corresponding to the full length genes, the labeled probe sequence is based on the polynucleotide sequences of the invention. Random primers or primers specific to the library vector can be used to amplify the trapped cDNA. Such gene trapping techniques are described in Gruber et al., WO 95/04745 and Gruber et al., USPN 5,500,356. Kits are commercially available to perform gene trapping experiments from, for example, Life Technologies, Gaithersburg, Maryland, USA.

"Rapid amplification of cDNA ends," or RACE, is a PCR method of amplifying cDNAs from a number of different RNAs. The cDNAs are ligated to an oligonucleotide linker, and amplified by PCR using two primers. One primer is based on sequence from the instant polynucleotides, for which full length sequence is desired, and a second primer comprises sequence that hybridizes to the oligonucleotide linker to amplify the cDNA. A description of this method is reported in WO 97/19110. In preferred embodiments of RACE, a common primer is designed to anneal to an arbitrary adaptor sequence ligated to cDNA ends (Apte and Siebert, *Biotechniques* (1993) 15:890-893; Edwards et al., *Nuc. Acids Res.* (1991) 19:5227-5232). When a single gene-specific RACE primer is paired with the common primer, preferential amplification of sequences between the single gene specific primer and the common primer occurs. Commercial cDNA pools modified for use in RACE are available.

Another PCR-based method generates full-length cDNA library with anchored ends without needing specific knowledge of the cDNA sequence. The method uses lock-docking primers (I-VI), where one primer, poly TV (I-III) locks over the polyA tail of eukaryotic mRNA producing first strand synthesis and a second primer, polyGH (IV-VI) locks onto the polyC tail added by terminal deoxynucleotidyl transferase (TdT)(see, e.g., WO 96/40998).

The promoter region of a gene generally is located 5' to the initiation site for RNA polymerase II. Hundreds of promoter regions contain the "TATA" box, a sequence such as TATTA or TATAA, which is sensitive to mutations. The promoter region can be obtained by performing 5' RACE using a primer from the coding region of the gene. Alternatively, the cDNA can be used as a probe for the genomic sequence, and the region 5' to the coding region is identified by "walking up." If the gene is highly expressed or differentially expressed, the promoter from the gene can be of use in a regulatory construct for a heterologous gene.

Once the full-length cDNA or gene is obtained, DNA encoding variants can be prepared by site-directed mutagenesis, described in detail in Sambrook et al., 15.3-15.63. The choice of codon or

nucleotide to be replaced can be based on disclosure herein on optional changes in amino acids to achieve altered protein structure and/or function.

As an alternative method to obtaining DNA or RNA from a biological material, nucleic acid comprising nucleotides having the sequence of one or more polynucleotides of the invention can be synthesized. Thus, the invention encompasses nucleic acid molecules ranging in length from 15 nt (corresponding to at least 15 contiguous nt of one of SEQ ID NOS:1-1485) up to a maximum length suitable for one or more biological manipulations, including replication and expression, of the nucleic acid molecule. The invention includes but is not limited to (a) nucleic acid having the size of a full gene, and comprising at least one of SEQ ID NOS:1-1485; (b) the nucleic acid of (a) also comprising at least one additional gene, operably linked to permit expression of a fusion protein; (c) an expression vector comprising (a) or (b); (d) a plasmid comprising (a) or (b); and (e) a recombinant viral particle comprising (a) or (b). Once provided with the polynucleotides disclosed herein, construction or preparation of (a) - (e) are well within the skill in the art.

The sequence of a nucleic acid comprising at least 15 contiguous nt of at least any one of SEQ ID NOS:1-1485, preferably the entire sequence of at least any one of SEQ ID NOS:1-1485, is not limited and can be any sequence of A, T, G, and/or C (for DNA) and A, U, G, and/or C (for RNA) or modified bases thereof, including inosine and pseudouridine. The choice of sequence will depend on the desired function and can be dictated by coding regions desired, the intron-like regions desired, and the regulatory regions desired. Where the entire sequence of any one of SEQ ID NOS:1-1485 is within the nucleic acid, the nucleic acid obtained is referred to herein as a polynucleotide comprising the sequence of any one of SEQ ID NOS:1-1485.

#### Expression of Polypeptide Encoded by Full-Length cDNA or Full-Length Gene

The provided polynucleotides (e.g., a polynucleotide having a sequence of one of SEQ ID NOS:1-1485), the corresponding cDNA, or the full-length gene is used to express a partial or complete gene product. Constructs of polynucleotides having sequences of SEQ ID NOS:1-1485 can also be generated synthetically. Alternatively, single-step assembly of a gene and entire plasmid from large numbers of oligodeoxyribonucleotides is described by, e.g., Stemmer et al., *Gene* (Amsterdam) (1995) 164(1):49-53. In this method, assembly PCR (the synthesis of long DNA sequences from large numbers of oligodeoxyribonucleotides (oligos)) is described. The method is derived from DNA shuffling (Stemmer, *Nature* (1994) 370:389-391), and does not rely on DNA ligase, but instead relies on DNA polymerase to build increasingly longer DNA fragments during the assembly process.

Appropriate polynucleotide constructs are purified using standard recombinant DNA techniques as described in, for example, Sambrook et al., *Molecular Cloning: A Laboratory Manual*, 2nd Ed., (1989) Cold Spring Harbor Press, Cold Spring Harbor, NY, and under current regulations described in United States Dept. of HHS, National Institute of Health (NIH) Guidelines for Recombinant DNA Research. The gene product encoded by a polynucleotide of the invention is

expressed in any expression system, including, for example, bacterial, yeast, insect, amphibian and mammalian systems. Vectors, host cells and methods for obtaining expression in same are well known in the art. Suitable vectors and host cells are described in USPN 5,654,173.

Polynucleotide molecules comprising a polynucleotide sequence provided herein are generally propagated by placing the molecule in a vector. Viral and non-viral vectors are used, including plasmids. The choice of plasmid will depend on the type of cell in which propagation is desired and the purpose of propagation. Certain vectors are useful for amplifying and making large amounts of the desired DNA sequence. Other vectors are suitable for expression in cells in culture. Still other vectors are suitable for transfer and expression in cells in a whole animal or person. The choice of appropriate vector is well within the skill of the art. Many such vectors are available commercially. Methods for preparation of vectors comprising a desired sequence are well known in the art.

The polynucleotides set forth in SEQ ID NOS:1-1485 or their corresponding full-length polynucleotides are linked to regulatory sequences as appropriate to obtain the desired expression properties. These can include promoters (attached either at the 5' end of the sense strand or at the 3' end of the antisense strand), enhancers, terminators, operators, repressors, and inducers. The promoters can be regulated or constitutive. In some situations it may be desirable to use conditionally active promoters, such as tissue-specific or developmental stage-specific promoters. These are linked to the desired nucleotide sequence using the techniques described above for linkage to vectors. Any techniques known in the art can be used.

When any of the above host cells, or other appropriate host cells or organisms, are used to replicate and/or express the polynucleotides or nucleic acids of the invention, the resulting replicated nucleic acid, RNA, expressed protein or polypeptide, is within the scope of the invention as a product of the host cell or organism. The product is recovered by any appropriate means known in the art.

Once the gene corresponding to a selected polynucleotide is identified, its expression can be regulated in the cell to which the gene is native. For example, an endogenous gene of a cell can be regulated by an exogenous regulatory sequence as disclosed in USPN 5,641,670.

#### Identification of Functional and Structural Motifs

Translations of the nucleotide sequence of the provided polynucleotides, cDNAs or full genes can be aligned with individual known sequences. Similarity with individual sequences can be used to determine the activity of the polypeptides encoded by the polynucleotides of the invention. Also, sequences exhibiting similarity with more than one individual sequence can exhibit activities that are characteristic of either or both individual sequences.

The full length sequences and fragments of the polynucleotide sequences of the nearest neighbors as identified through, for example, BLAST-based searching, can be used as probes and primers to identify and isolate the full length sequence corresponding to provided polynucleotides.

The nearest neighbors can indicate a tissue or cell type to be used to construct a library for the full-length sequences corresponding to the provided polynucleotides.

Typically, a selected polynucleotide is translated in all six frames to determine the best alignment with the individual sequences. The sequences disclosed herein in the Sequence Listing are in a 5' to 3' orientation and translation in three frames can be sufficient (with a few specific exceptions as described in the Examples). These amino acid sequences are referred to, generally, as query sequences, which will be aligned with the individual sequences. Databases with individual sequences are described in "Computer Methods for Macromolecular Sequence Analysis" *Methods in Enzymology* (1996) 266, Doolittle, Academic Press, Inc., a division of Harcourt Brace & Co., San Diego, California, USA. Databases include GenBank, EMBL, and DNA Database of Japan (DDBJ).

Query and individual sequences can be aligned using the methods and computer programs described above, and include BLAST 2.0, available over the world wide web at a site supported by the National Center for Biotechnology Information, which is supported by the National Library of Medicine and the National Institutes of Health, or TeraBLAST available from TimeLogic Corp. (Crystal Bay, Nevada). See also Altschul, et al. *Nucleic Acids Res.* (1997) 25:3389-3402. Another alignment algorithm is Fasta, available in the Genetics Computing Group (GCG) package, Madison, Wisconsin, USA, a wholly owned subsidiary of Oxford Molecular Group, Inc. Other techniques for alignment are described in Doolittle, *supra*. Preferably, an alignment program that permits gaps in the sequence is utilized to align the sequences. The Smith-Waterman is one type of algorithm that permits gaps in sequence alignments. See *Meth. Mol. Biol.* (1997) 70: 173-187. Also, the GAP program using the Needleman and Wunsch alignment method can be utilized to align sequences. An alternative search strategy uses MPSRCH software, which runs on a MASPAR computer. MPSRCH uses a Smith-Waterman algorithm to score sequences on a massively parallel computer. This approach improves ability to identify sequences that are distantly related matches, and is especially tolerant of small gaps and nucleotide sequence errors. Amino acid sequences encoded by the provided polynucleotides can be used to search both protein and DNA databases. Incorporated herein by reference are all sequences that have been made public as of the filing date of this application by any of the DNA or protein sequence databases, including the patent databases (e.g., GeneSeq). Also incorporated by reference are those sequences that have been submitted to these databases as of the filing date of the present application but not made public until after the filing date of the present application.

Results of individual and query sequence alignments can be divided into three categories: high similarity, weak similarity, and no similarity. Individual alignment results ranging from high similarity to weak similarity provide a basis for determining polypeptide activity and/or structure. Parameters for categorizing individual results include: percentage of the alignment region length where the strongest alignment is found, percent sequence identity, and p value. The percentage of the

alignment region length is calculated by counting the number of residues of the individual sequence found in the region of strongest alignment, e.g., contiguous region of the individual sequence that contains the greatest number of residues that are identical to the residues of the corresponding region of the aligned query sequence. This number is divided by the total residue length of the query sequence to calculate a percentage. For example, a query sequence of 20 amino acid residues might be aligned with a 20 amino acid region of an individual sequence. The individual sequence might be identical to amino acid residues 5, 9-15, and 17-19 of the query sequence. The region of strongest alignment is thus the region stretching from residue 9-19, an 11 amino acid stretch. The percentage of the alignment region length is: 11 (length of the region of strongest alignment) divided by (query sequence length) 20 or 55%.

Percent sequence identity is calculated by counting the number of amino acid matches between the query and individual sequence and dividing total number of matches by the number of residues of the individual sequences found in the region of strongest alignment. Thus, the percent identity in the example above would be 10 matches divided by 11 amino acids, or approximately, 90.9%.

P value is the probability that the alignment was produced by chance. For a single alignment, the p value can be calculated according to Karlin et al., Proc. Natl. Acad. Sci. (1990) 87:2264 and Karlin et al., Proc. Natl. Acad. Sci. (1993) 90. The p value of multiple alignments using the same query sequence can be calculated using an heuristic approach described in Altschul et al., Nat. Genet. (1994) 6:119. Alignment programs, such as BLAST or TeraBLAST, can calculate the p value. See also Altschul et al., Nucleic Acids Res. (1997) 25:3389-3402.

Another factor to consider for determining identity or similarity is the location of the similarity or identity. Strong local alignment can indicate similarity even if the length of alignment is short. Sequence identity scattered throughout the length of the query sequence also can indicate a similarity between the query and profile sequences. The boundaries of the region where the sequences align can be determined according to Doolittle, supra; BLAST 2.0 (see, e.g., Altschul, et al. Nucleic Acids Res. (1997) 25:3389-3402), TeraBLAST (available from TimeLogic Corp., Crystal Bay, Nevada), or FAST programs; or by determining the area where sequence identity is highest.

High Similarity. In general, in alignment results considered to be of high similarity, the percent of the alignment region length is typically at least about 55% of total length query sequence; more typically, at least about 58%; even more typically, at least about 60% of the total residue length of the query sequence. Usually, percent length of the alignment region can be as much as about 62%; more usually, as much as about 64%; even more usually, as much as about 66%. Further, for high similarity, the region of alignment, typically, exhibits at least about 75% of sequence identity; more typically, at least about 78%; even more typically, at least about 80% sequence identity. Usually,

percent sequence identity can be as much as about 82%; more usually, as much as about 84%; even more usually, as much as about 86%.

The p value is used in conjunction with these methods. If high similarity is found, the query sequence is considered to have high similarity with a profile sequence when the p value is less than or equal to about  $10e-2$ ; more usually, less than or equal to about  $10e-3$ ; even more usually, less than or equal to about  $10e-4$ . More typically, the p value is no more than about  $10e-5$ ; more typically, no more than or equal to about  $10e-10$ ; even more typically, no more than or equal to about  $10e-15$  for the query sequence to be considered high similarity.

Weak Similarity. In general, where alignment results considered to be of weak similarity, there is no minimum percent length of the alignment region nor minimum length of alignment. A better showing of weak similarity is considered when the region of alignment is, typically, at least about 15 amino acid residues in length; more typically, at least about 20; even more typically, at least about 25 amino acid residues in length. Usually, length of the alignment region can be as much as about 30 amino acid residues; more usually, as much as about 40; even more usually, as much as about 60 amino acid residues. Further, for weak similarity, the region of alignment, typically, exhibits at least about 35% of sequence identity; more typically, at least about 40%; even more typically, at least about 45% sequence identity. Usually, percent sequence identity can be as much as about 50%; more usually, as much as about 55%; even more usually, as much as about 60%.

If low similarity is found, the query sequence is considered to have weak similarity with a profile sequence when the p value is usually less than or equal to about  $10e-2$ ; more usually, less than or equal to about  $10e-3$ ; even more usually, less than or equal to about  $10e-4$ . More typically, the p value is no more than about  $10e-5$ ; more usually, no more than or equal to about  $10e-10$ ; even more usually, no more than or equal to about  $10e-15$  for the query sequence to be considered weak similarity.

Similarity Determined by Sequence Identity Alone. Sequence identity alone can be used to determine similarity of a query sequence to an individual sequence and can indicate the activity of the sequence. Such an alignment, preferably, permits gaps to align sequences. Typically, the query sequence is related to the profile sequence if the sequence identity over the entire query sequence is at least about 15%; more typically, at least about 20%; even more typically, at least about 25%; even more typically, at least about 50%. Sequence identity alone as a measure of similarity is most useful when the query sequence is usually, at least 80 residues in length; more usually, at least 90 residues in length; even more usually, at least 95 amino acid residues in length. More typically, similarity can be concluded based on sequence identity alone when the query sequence is preferably 100 residues in length; more preferably, 120 residues in length; even more preferably, 150 amino acid residues in length.



Alignments with Profile and Multiple Aligned Sequences. Translations of the provided polynucleotides can be aligned with amino acid profiles that define either protein families or common motifs. Also, translations of the provided polynucleotides can be aligned to multiple sequence alignments (MSA) comprising the polypeptide sequences of members of protein families or motifs.

5 Similarity or identity with profile sequences or MSAs can be used to determine the activity of the gene products (e.g., polypeptides) encoded by the provided polynucleotides or corresponding cDNA or genes. For example, sequences that show an identity or similarity with a chemokine profile or MSA can exhibit chemokine activities.

Profiles can be designed manually by (1) creating an MSA, which is an alignment of the amino acid sequence of members that belong to the family and (2) constructing a statistical representation of the alignment. Such methods are described, for example, in Birney et al., Nucl. Acid Res. (1996) 24(14): 2730-2739. MSAs of some protein families and motifs are publicly available. For example, the Genome Sequencing Center at the Washington University School of Medicine provides a web set (Pfam) which provides MSAs of 547 different families and motifs. These MSAs are described also in Sonnhammer et al., Proteins (1997) 28: 405-420. Other sources over the world wide web include the site supported by the European Molecular Biology Laboratories in Heidelberg, Germany. A brief description of these MSAs is reported in Pascarella et al., Prot. Eng. (1996) 9(3):249-251. Techniques for building profiles from MSAs are described in Sonnhammer et al., supra; Birney et al., supra; and "Computer Methods for Macromolecular Sequence Analysis," Methods in

20 Enzymology (1996) 266, Doolittle, Academic Press, Inc., San Diego, California, USA.

Similarity between a query sequence and a protein family or motif can be determined by (a) comparing the query sequence against the profile and/or (b) aligning the query sequence with the members of the family or motif. Typically, a program such as Searchwise is used to compare the query sequence to the statistical representation of the multiple alignment, also known as a profile (see

25 Birney et al., supra). Other techniques to compare the sequence and profile are described in Sonnhammer et al., supra and Doolittle, supra.

Next, methods described by Feng et al., J. Mol. Evol. (1987) 25:351 and Higgins et al., CABIOS (1989) 5:151 can be used align the query sequence with the members of a family or motif, also known as a MSA. Sequence alignments can be generated using any of a variety of software tools.

30 Examples include PileUp, which creates a multiple sequence alignment, and is described in Feng et al., J. Mol. Evol. (1987) 25:351. Another method, GAP, uses the alignment method of Needleman et al., J. Mol. Biol. (1970) 48:443. GAP is best suited for global alignment of sequences. A third method, BestFit, functions by inserting gaps to maximize the number of matches using the local homology algorithm of Smith et al., Adv. Appl. Math. (1981) 2:482. In general, the following factors

35 are used to determine if a similarity between a query sequence and a profile or MSA exists: (1)

number of conserved residues found in the query sequence, (2) percentage of conserved residues found in the query sequence, (3) number of frameshifts, and (4) spacing between conserved residues.

Some alignment programs that both translate and align sequences can make any number of frameshifts when translating the nucleotide sequence to produce the best alignment. The fewer frameshifts needed to produce an alignment, the stronger the similarity or identity between the query and profile or MSAs. For example, a weak similarity resulting from no frameshifts can be a better indication of activity or structure of a query sequence, than a strong similarity resulting from two frameshifts. Preferably, three or fewer frameshifts are found in an alignment; more preferably two or fewer frameshifts; even more preferably, one or fewer frameshifts; even more preferably, no frameshifts are found in an alignment of query and profile or MSAs.

Conserved residues are those amino acids found at a particular position in all or some of the family or motif members. Alternatively, a position is considered conserved if only a certain class of amino acids is found in a particular position in all or some of the family members. For example, the N-terminal position can contain a positively charged amino acid, such as lysine, arginine, or histidine.

Typically, a residue of a polypeptide is conserved when a class of amino acids or a single amino acid is found at a particular position in at least about 40% of all class members; more typically, at least about 50%; even more typically, at least about 60% of the members. Usually, a residue is conserved when a class or single amino acid is found in at least about 70% of the members of a family or motif; more usually, at least about 80%; even more usually, at least about 90%; even more usually, at least about 95%.

A residue is considered conserved when three unrelated amino acids are found at a particular position in some or all of the members; more usually, two unrelated amino acids. These residues are conserved when the unrelated amino acids are found at particular positions in at least about 40% of all class member; more typically, at least about 50%; even more typically, at least about 60% of the members. Usually, a residue is conserved when a class or single amino acid is found in at least about 70% of the members of a family or motif; more usually, at least about 80%; even more usually, at least about 90%; even more usually, at least about 95%.

A query sequence has similarity to a profile or MSA when the query sequence comprises at least about 25% of the conserved residues of the profile or MSA; more usually, at least about 30%; even more usually; at least about 40%. Typically, the query sequence has a stronger similarity to a profile sequence or MSA when the query sequence comprises at least about 45% of the conserved residues of the profile or MSA; more typically, at least about 50%; even more typically, at least about 55%.

Identification of Secreted & Membrane-Bound Polypeptides. Both secreted and membrane-bound polypeptides of the present invention are of particular interest. For example, levels of secreted polypeptides can be assayed in body fluids that are convenient, such as blood, plasma, serum, and

other body fluids such as urine, prostatic fluid and semen. Membrane-bound polypeptides are useful for constructing vaccine antigens or inducing an immune response. Such antigens would comprise all or part of the extracellular region of the membrane-bound polypeptides. Because both secreted and membrane-bound polypeptides comprise a fragment of contiguous hydrophobic amino acids, hydrophobicity predicting algorithms can be used to identify such polypeptides.

A signal sequence is usually encoded by both secreted and membrane-bound polypeptide genes to direct a polypeptide to the surface of the cell. The signal sequence usually comprises a stretch of hydrophobic residues. Such signal sequences can fold into helical structures. Membrane-bound polypeptides typically comprise at least one transmembrane region that possesses a stretch of hydrophobic amino acids that can transverse the membrane. Some transmembrane regions also exhibit a helical structure. Hydrophobic fragments within a polypeptide can be identified by using computer algorithms. Such algorithms include Hopp & Woods, Proc. Natl. Acad. Sci. USA (1981) 78:3824-3828; Kyte & Doolittle, J. Mol. Biol. (1982) 157: 105-132; and RAOAR algorithm, Degli Esposti et al., Eur. J. Biochem. (1990) 190: 207-219.

Another method of identifying secreted and membrane-bound polypeptides is to translate the polynucleotides of the invention in all six frames and determine if at least 8 contiguous hydrophobic amino acids are present. Those translated polypeptides with at least 8; more typically, 10; even more typically, 12 contiguous hydrophobic amino acids are considered to be either a putative secreted or membrane bound polypeptide. Hydrophobic amino acids include alanine, glycine, histidine, isoleucine, leucine, lysine, methionine, phenylalanine, proline, threonine, tryptophan, tyrosine, and valine

#### Identification of the Function of an Expression Product of a Full-Length Gene

Ribozymes, antisense constructs, and dominant negative mutants can be used to determine function of the expression product of a gene corresponding to a polynucleotide provided herein.

These methods and compositions are particularly useful where the provided novel polynucleotide exhibits no significant or substantial homology to a sequence encoding a gene of known function. Antisense molecules and ribozymes can be constructed from synthetic polynucleotides. Typically, the phosphoramidite method of oligonucleotide synthesis is used. See Beaucage et al., Tet. Lett. (1981) 22:1859 and USPN 4,668,777. Automated devices for synthesis are available to create oligonucleotides using this chemistry. Examples of such devices include Biosearch 8600, Models 392 and 394 by Applied Biosystems, a division of Perkin-Elmer Corp., Foster City, California, USA; and Expedite by Perceptive Biosystems, Framingham, Massachusetts, USA. Synthetic RNA, phosphate analog oligonucleotides, and chemically derivatized oligonucleotides can also be produced, and can be covalently attached to other molecules. RNA oligonucleotides can be synthesized, for example, using RNA phosphoramidites. This method can be performed on an automated synthesizer, such as Applied Biosystems, Models 392 and 394, Foster City, California, USA.

Phosphorothioate oligonucleotides can also be synthesized for antisense construction. A sulfurizing reagent, such as tetraethylthiuram disulfide (TETD) in acetonitrile can be used to convert the internucleotide cyanoethyl phosphite to the phosphorothioate triester within 15 minutes at room temperature. TETD replaces the iodine reagent, while all other reagents used for standard phosphoramidite chemistry remain the same. Such a synthesis method can be automated using Models 392 and 394 by Applied Biosystems, for example.

Oligonucleotides of up to 200 nt can be synthesized, more typically, 100 nt; more typically 50 nt; even more typically, 30 to 40 nt. These synthetic fragments can be annealed and ligated together to construct larger fragments. See, for example, Sambrook et al., *supra*. Trans-cleaving catalytic RNAs (ribozymes) are RNA molecules possessing endoribonuclease activity. Ribozymes are specifically designed for a particular target, and the target message must contain a specific nucleotide sequence. They are engineered to cleave any RNA species site-specifically in the background of cellular RNA. The cleavage event renders the mRNA unstable and prevents protein expression. Importantly, ribozymes can be used to inhibit expression of a gene of unknown function for the purpose of determining its function in an in vitro or in vivo context, by detecting the phenotypic effect. One commonly used ribozyme motif is the hammerhead, for which the substrate sequence requirements are minimal. Design of the hammerhead ribozyme, as well as therapeutic uses of ribozymes, are disclosed in Usman et al., *Current Opin. Struct. Biol.* (1996) 6:527. Methods for production of ribozymes, including hairpin structure ribozyme fragments, methods of increasing ribozyme specificity, and the like are known in the art.

The hybridizing region of the ribozyme can be modified or can be prepared as a branched structure as described in Horn and Urdea, *Nucleic Acids Res.* (1989) 17:6959. The basic structure of the ribozymes can also be chemically altered in ways familiar to those skilled in the art, and chemically synthesized ribozymes can be administered as synthetic oligonucleotide derivatives modified by monomeric units. In a therapeutic context, liposome mediated delivery of ribozymes improves cellular uptake, as described in Birikh et al., *Eur. J. Biochem.* (1997) 245:1.

Antisense nucleic acids are designed to specifically bind to RNA, resulting in the formation of RNA-DNA or RNA-RNA hybrids, with an arrest of DNA replication, reverse transcription or messenger RNA translation. Antisense polynucleotides based on a selected polynucleotide sequence can interfere with expression of the corresponding gene. Antisense polynucleotides are typically generated within the cell by expression from antisense constructs that contain the antisense strand as the transcribed strand. Antisense polynucleotides based on the disclosed polynucleotides will bind and/or interfere with the translation of mRNA comprising a sequence complementary to the antisense polynucleotide. The expression products of control cells and cells treated with the antisense construct are compared to detect the protein product of the gene corresponding to the polynucleotide upon

which the antisense construct is based. The protein is isolated and identified using routine biochemical methods.

Given the extensive background literature and clinical experience in antisense therapy, one skilled in the art can use selected polynucleotides of the invention as additional potential therapeutics.

- 5 The choice of polynucleotide can be narrowed by first testing them for binding to "hot spot" regions of the genome of cancerous cells. If a polynucleotide is identified as binding to a "hot spot," testing the polynucleotide as an antisense compound in the corresponding cancer cells is warranted.

- As an alternative method for identifying function of the gene corresponding to a polynucleotide disclosed herein, dominant negative mutations are readily generated for corresponding proteins that are active as homomultimers. A mutant polypeptide will interact with wild-type polypeptides (made from the other allele) and form a non-functional multimer. Thus, a mutation is in a substrate-binding domain, a catalytic domain, or a cellular localization domain. Preferably, the mutant polypeptide will be overproduced. Point mutations are made that have such an effect. In addition, fusion of different polypeptides of various lengths to the terminus of a protein can yield dominant negative mutants. General strategies are available for making dominant negative mutants (see, e.g., Herskowitz, Nature (1987) 329:219). Such techniques can be used to create loss of function mutations, which are useful for determining protein function.
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#### Polypeptides and Variants Thereof

- The polypeptides of the invention include those encoded by the disclosed polynucleotides, as well as nucleic acids that, by virtue of the degeneracy of the genetic code, are not identical in sequence to the disclosed polynucleotides. Thus, the invention includes within its scope a polypeptide encoded by a polynucleotide having the sequence of any one of SEQ ID NOS:1-1485 or a variant thereof. Also included in the invention are the polypeptides comprising the amino acid sequences of SEQ ID NOS:1486-1542.
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- In general, the term "polypeptide" as used herein refers to both the full length polypeptide encoded by the recited polynucleotide, the polypeptide encoded by the gene represented by the recited polynucleotide, as well as portions or fragments thereof. "Polypeptides" also includes variants of the naturally occurring proteins, where such variants are homologous or substantially similar to the naturally occurring protein, and can be of an origin of the same or different species as the naturally occurring protein (e.g., human, murine, or some other species that naturally expresses the recited polypeptide, usually a mammalian species). In general, variant polypeptides have a sequence that has at least about 80%, usually at least about 90%, and more usually at least about 98% sequence identity with a differentially expressed polypeptide of the invention, as measured by BLAST 2.0 or TeraBLAST using the parameters described above. The variant polypeptides can be naturally or non-naturally glycosylated, i.e., the polypeptide has a glycosylation pattern that differs from the glycosylation pattern found in the corresponding naturally occurring protein.
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The invention also encompasses homologs of the disclosed polypeptides (or fragments thereof) where the homologs are isolated from other species, i.e. other animal or plant species, where such homologs, usually mammalian species, e.g. rodents, such as mice, rats; domestic animals, e.g., horse, cow, dog, cat; and humans. By "homolog" is meant a polypeptide having at least about 35%, usually at least about 40% and more usually at least about 60% amino acid sequence identity to a particular differentially expressed protein as identified above, where sequence identity is determined using the BLAST 2.0 or TeraBLAST algorithm, with the parameters described supra.

In general, the polypeptides of the subject invention are provided in a non-naturally occurring environment, e.g. are separated from their naturally occurring environment. In certain embodiments, the subject protein is present in a composition that is enriched for the protein as compared to a control. As such, purified polypeptide is provided, where by purified is meant that the protein is present in a composition that is substantially free of non-differentially expressed polypeptides, where by substantially free is meant that less than 90%, usually less than 60% and more usually less than 50% of the composition is made up of non-differentially expressed polypeptides.

Also within the scope of the invention are variants; variants of polypeptides include mutants, fragments, and fusions. Mutants can include amino acid substitutions, additions or deletions. The amino acid substitutions can be conservative amino acid substitutions or substitutions to eliminate non-essential amino acids, such as to alter a glycosylation site, a phosphorylation site or an acetylation site, or to minimize misfolding by substitution or deletion of one or more cysteine residues that are not necessary for function. Conservative amino acid substitutions are those that preserve the general charge, hydrophobicity/ hydrophilicity, and/or steric bulk of the amino acid substituted. Variants can be designed so as to retain or have enhanced biological activity of a particular region of the protein (e.g., a functional domain and/or, where the polypeptide is a member of a protein family, a region associated with a consensus sequence). Selection of amino acid alterations for production of variants can be based upon the accessibility (interior vs. exterior) of the amino acid (see, e.g., Go et al, *Int. J. Peptide Protein Res.* (1980) 15:211), the thermostability of the variant polypeptide (see, e.g., Querol et al., *Prot. Eng.* (1996) 9:265), desired glycosylation sites (see, e.g., Olsen and Thomsen, *J. Gen. Microbiol.* (1991) 137:579), desired disulfide bridges (see, e.g., Clarke et al., *Biochemistry* (1993) 32:4322; and Wakarchuk et al., *Protein Eng.* (1994) 7:1379), desired metal binding sites (see, e.g., Toma et al., *Biochemistry* (1991) 30:97, and Haezzebrouck et al., *Protein Eng.* (1993) 6:643), and desired substitutions within proline loops (see, e.g., Masul et al., *Appl. Env. Microbiol.* (1994) 60:3579). Cysteine-depleted muteins can be produced as disclosed in USPN 4,959,314.

Variants also include fragments of the polypeptides disclosed herein, particularly haptens, biologically active fragments, and/or fragments corresponding to functional domains. Fragments of interest will typically be at least about 10 aa to at least about 15 aa in length, usually at least about 50 aa in length, and can be as long as 300 aa in length or longer, but will usually not exceed about 1000

aa in length, where the fragment will have a stretch of amino acids that is identical to a polypeptide encoded by a polynucleotide having a sequence of any SEQ ID NOS:1-1485, a polypeptide comprising a sequence of at least one of SEQ ID NOS:1486-1542, or a homolog thereof. The protein variants described herein are encoded by polynucleotides that are within the scope of the invention.

- 5 The genetic code can be used to select the appropriate codons to construct the corresponding variants.

A fragment of a subject polypeptide is, for example, a polypeptide having an amino acid sequence which is a portion of a subject polypeptide e.g. a polypeptide encoded by a subject polynucleotide that is identified by any one of the sequence the sequence listing or its complement. The polypeptide fragments of the invention are preferably at least about 9 aa, at least about 15 aa, and more preferably at least about 20 aa, still more preferably at least about 30 aa, and even more preferably, at least about 40 aa, at least about 50 aa, at least about 75 aa, at least about 100 aa, at least about 125 aa or at least about 150 aa in length. A fragment "at least 20 aa in length," for example, is intended to include 20 or more contiguous amino acids from, for example, the polypeptide encoded by a cDNA, in a cDNA clone contained in a deposited library, or a nucleotide sequence shown in the sequence listing or the complementary stand thereof. In this context "about" includes the particularly recited value or a value larger or smaller by several (5, 4, 3, 2, or 1) amino acids. These polypeptide fragments have uses that include, but are not limited to, production of antibodies as discussed herein. Of course, larger fragments (e.g., at least 150, 175, 200, 250, 500, 600, 1000, or 2000 amino acids in length) are also encompassed by the invention.

- 20 Moreover, representative examples of polypeptides fragments of the invention (useful in, for example, as antigens for antibody production), include, for example, fragments comprising, or alternatively consisting of, a sequence from about amino acid number 1-10, 5-10, 10-20, 21-31, 31-40, 41-61, 61-81, 91-120, 121-140, 141-162, 162-200, 201-240, 241-280, 281- 320, 321-360, 360-400, 400-450, 451-500, 500-600, 600-700, 700-800, 800-900 and the like. In this context "about" includes the particularly recited range or a range larger or smaller by several (5, 4, 3, 2, or 1) amino acids, at either terminus or at both termini. In some embodiments, these fragments has a functional activity (e.g., biological activity) whereas in other embodiments, these fragments may be used to make an antibody.

- 30 Further polypeptide variants may are described in PCT publications WO/00-55173, WO/01-07611 and WO/02-16429

#### Computer-Related Embodiments

- In general, a library of polynucleotides is a collection of sequence information, which information is provided in either biochemical form (e.g., as a collection of polynucleotide molecules), or in electronic form (e.g., as a collection of polynucleotide sequences stored in a computer-readable form, as in a computer system and/or as part of a computer program). The sequence information of the polynucleotides can be used in a variety of ways, e.g., as a resource for gene discovery, as a

representation of sequences expressed in a selected cell type (e.g., cell type markers), and/or as markers of a given disease or disease state. In general, a disease marker is a representation of a gene product that is present in all cells affected by disease either at an increased or decreased level relative to a normal cell (e.g., a cell of the same or similar type that is not substantially affected by disease).

- 5 For example, a polynucleotide sequence in a library can be a polynucleotide that represents an mRNA, polypeptide, or other gene product encoded by the polynucleotide, that is either overexpressed or underexpressed in a breast ductal cell affected by cancer relative to a normal (i.e., substantially disease-free) breast cell.

- The nucleotide sequence information of the library can be embodied in any suitable form, e.g., electronic or biochemical forms. For example, a library of sequence information embodied in electronic form comprises an accessible computer data file (or, in biochemical form, a collection of nucleic acid molecules) that contains the representative nucleotide sequences of genes that are differentially expressed (e.g., overexpressed or underexpressed) as between, for example, i) a cancerous cell and a normal cell; ii) a cancerous cell and a dysplastic cell; iii) a cancerous cell and a cell affected by a disease or condition other than cancer; iv) a metastatic cancerous cell and a normal cell and/or non-metastatic cancerous cell; v) a malignant cancerous cell and a non-malignant cancerous cell (or a normal cell) and/or vi) a dysplastic cell relative to a normal cell. Other combinations and comparisons of cells affected by various diseases or stages of disease will be readily apparent to the ordinarily skilled artisan. Biochemical embodiments of the library include a collection of nucleic acids that have the sequences of the genes in the library, where the nucleic acids can correspond to the entire gene in the library or to a fragment thereof, as described in greater detail below.

- The polynucleotide libraries of the subject invention generally comprise sequence information of a plurality of polynucleotide sequences, where at least one of the polynucleotides has a sequence of any of SEQ ID NOS:1-1485. By plurality is meant at least 2, usually at least 3 and can include up to all of SEQ ID NOS:1-1485. The length and number of polynucleotides in the library will vary with the nature of the library, e.g., if the library is an oligonucleotide array, a cDNA array, a computer database of the sequence information, etc.

- Where the library is an electronic library, the nucleic acid sequence information can be present in a variety of media. "Media" refers to a manufacture, other than an isolated nucleic acid molecule, that contains the sequence information of the present invention. Such a manufacture provides the genome sequence or a subset thereof in a form that can be examined by means not directly applicable to the sequence as it exists in a nucleic acid. For example, the nucleotide sequence of the present invention, e.g. the nucleic acid sequences of any of the polynucleotides of SEQ ID NOS:1-1485, can be recorded on computer readable media, e.g. any medium that can be read and accessed directly by a computer. Such media include, but are not limited to: magnetic storage media,



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such as a floppy disc, a hard disc storage medium, and a magnetic tape; optical storage media such as CD-ROM; electrical storage media such as RAM and ROM; and hybrids of these categories such as magnetic/optical storage media. One of skill in the art can readily appreciate how any of the presently known computer readable mediums can be used to create a manufacture comprising a recording of the present sequence information. "Recorded" refers to a process for storing information on computer readable medium, using any such methods as known in the art. Any convenient data storage structure can be chosen, based on the means used to access the stored information. A variety of data processor programs and formats can be used for storage, e.g. word processing text file, database format, etc. In addition to the sequence information, electronic versions of the libraries of the invention can be provided in conjunction or connection with other computer-readable information and/or other types of computer-readable files (e.g., searchable files, executable files, etc, including, but not limited to, for example, search program software, etc.).

By providing the nucleotide sequence in computer readable form, the information can be accessed for a variety of purposes. Computer software to access sequence information is publicly available. For example, the gapped BLAST (Altschul et al. Nucleic Acids Res. (1997) 25:3389-3402) and BLAZE (Brutlag et al. Comp. Chem. (1993) 17:203) search algorithms on a Sybase system, or the TeraBLAST (TimeLogic, Crystal Bay, Nevada) program optionally running on a specialized computer platform available from TimeLogic, can be used to identify open reading frames (ORFs) within the genome that contain homology to ORFs from other organisms.

As used herein, "a computer-based system" refers to the hardware means, software means, and data storage means used to analyze the nucleotide sequence information of the present invention. The minimum hardware of the computer-based systems of the present invention comprises a central processing unit (CPU), input means, output means, and data storage means. A skilled artisan can readily appreciate that any one of the currently available computer-based system are suitable for use in the present invention. The data storage means can comprise any manufacture comprising a recording of the present sequence information as described above, or a memory access means that can access such a manufacture.

"Search means" refers to one or more programs implemented on the computer-based system, to compare a target sequence or target structural motif, or expression levels of a polynucleotide in a sample, with the stored sequence information. Search means can be used to identify fragments or regions of the genome that match a particular target sequence or target motif. A variety of known algorithms are publicly known and commercially available, e.g. MacPattern (EMBL), BLASTN and BLASTX (NCBI), TeraBLAST (TimeLogic, Crystal Bay, Nevada). A "target sequence" can be any polynucleotide or amino acid sequence of six or more contiguous nucleotides or two or more amino acids, preferably from about 10 to 100 amino acids or from about 30 to 300 nt. A variety of comparing means can be used to accomplish comparison of sequence information from a sample (e.g., to analyze

target sequences, target motifs, or relative expression levels) with the data storage means. A skilled artisan can readily recognize that any one of the publicly available homology search programs can be used as the search means for the computer based systems of the present invention to accomplish comparison of target sequences and motifs. Computer programs to analyze expression levels in a sample and in controls are also known in the art.

A "target structural motif," or "target motif," refers to any rationally selected sequence or combination of sequences in which the sequence(s) are chosen based on a three-dimensional configuration that is formed upon the folding of the target motif, or on consensus sequences of regulatory or active sites. There are a variety of target motifs known in the art. Protein target motifs include, but are not limited to, enzyme active sites and signal sequences. Nucleic acid target motifs include, but are not limited to, hairpin structures, promoter sequences and other expression elements such as binding sites for transcription factors.

A variety of structural formats for the input and output means can be used to input and output the information in the computer-based systems of the present invention. One format for an output means ranks the relative expression levels of different polynucleotides. Such presentation provides a skilled artisan with a ranking of relative expression levels to determine a gene expression profile.

As discussed above, the "library" of the invention also encompasses biochemical libraries of the polynucleotides of SEQ ID NOS:1-1485, e.g., collections of nucleic acids representing the provided polynucleotides. The biochemical libraries can take a variety of forms, e.g., a solution of cDNAs, a pattern of probe nucleic acids stably associated with a surface of a solid support (i.e., an array) and the like. Of particular interest are nucleic acid arrays in which one or more of SEQ ID NOS:1-1485 is represented on the array. By array is meant an article of manufacture that has at least a substrate with at least two distinct nucleic acid targets on one of its surfaces, where the number of distinct nucleic acids can be considerably higher, typically being at least 10, usually at least 20, and often at least 25 distinct nucleic acid molecules. A variety of different array formats have been developed and are known to those of skill in the art. The arrays of the subject invention find use in a variety of applications, including gene expression analysis, drug screening, mutation analysis and the like, as disclosed in the above-listed exemplary patent documents.

In addition to the above nucleic acid libraries, analogous libraries of polypeptides are also provided, where the polypeptides of the library will represent at least a portion of the polypeptides encoded by a gene corresponding to one or more of SEQ ID NOS:1-1485.

#### Utilities

The polynucleotides of the invention are useful in a variety of applications. Exemplary utilities of the polynucleotides of the invention are described below.

Construction of Larger Molecules: Recombinant DNAs and Nucleic Acid Multimers. In one embodiment of particular interest, the polynucleotides described herein as useful as the building

blocks for larger molecules. In one example, the polynucleotide is a component of a larger cDNA molecule which in turn can be adapted for expression in a host cell (e.g., a bacterial or eukaryotic (e.g., yeast or mammalian) host cell). The cDNA can include, in addition to the polypeptide encoded by the starting material polynucleotide (i.e., a polynucleotide described herein), an amino acid sequence that is heterologous to the polypeptide encoded by the polynucleotide described herein (e.g., as in a sequence encoding a fusion protein). In some embodiments, the polynucleotides described herein is used as starting material polynucleotide for synthesizing all or a portion of the gene to which the described polynucleotide corresponds. For example, a DNA molecule encoding a full-length human polypeptide can be constructed using a polynucleotide described herein as starting material.

In another embodiment, the polynucleotides of the invention are used in nucleic acid multimers. Nucleic acid multimers can be linear or branched polymers of the same repeating single-stranded oligonucleotide unit or different single-stranded oligonucleotide units. Where the molecules are branched, the multimers are generally described as either "fork" or "comb" structures. The oligonucleotide units of the multimer may be composed of RNA, DNA, modified nucleotides or combinations thereof. At least one of the units has a sequence, length, and composition that permits it to bind specifically to a first single-stranded nucleotide sequence of interest, typically analyte or an oligonucleotide bound to the analyte. In order to achieve such specificity and stability, this unit will normally be 15 to 50 nt, preferably 15 to 30 nt, in length and have a GC content in the range of 40% to 60%. In addition to such unit(s), the multimer includes a multiplicity of units that are capable of hybridizing specifically and stably to a second single-stranded nucleotide of interest, typically a labeled oligonucleotide or another multimer. These units will also normally be 15 to 50 nt, preferably 15 to 30 nt, in length and have a GC content in the range of 40% to 60%. When a multimer is designed to be hybridized to another multimer, the first and second oligonucleotide units are heterogeneous (different). One or more of the polynucleotides described herein, or a portion of a polynucleotide described herein, can be used as a repeating unit of such nucleic acid multimers.

The total number of oligonucleotide units in the multimer will usually be in the range of 3 to 50, more usually 10 to 20. In multimers in which the unit that hybridizes to the nucleotide sequence of interest is different from the unit that hybridizes to the labeled oligonucleotide, the number ratio of the latter to the former will usually be 2:1 to 30:1, more usually 5:1 to 20:1, and-preferably 10:1 to 15:1.

The oligonucleotide units of the multimer may be covalently linked directly to each other through phosphodiester bonds or through interposed linking agents such as nucleic acid, amino acid, carbohydrate or polyol bridges, or through other cross-linking agents that are capable of cross-linking nucleic acid or modified nucleic acid strands. The site(s) of linkage may be at the ends of the unit (in either normal 3',-5' orientation or randomly oriented) and/or at one or more internal nucleotides in the strand. In linear multimers the individual units are linked end-to-end to form a linear polymer. In one

type of branched multimer three or more oligonucleotide units emanate from a point of origin to form a branched structure. The point of origin may be another oligonucleotide unit or a multifunctional molecule to which at least three units can be covalently bound. In another type, there is an oligonucleotide unit backbone with one or more pendant oligonucleotide units. These latter-type multimers are "fork-like", "comb-like" or combination "fork-" and "comb-like" in structure. The pendant units will normally depend from a modified nucleotide or other organic moiety having appropriate functional groups to which oligonucleotides may be conjugated or otherwise attached. The multimer may be totally linear, totally branched, or a combination of linear and branched portions. Preferably there will be at least two branch points in the multimer, more preferably at least 3, preferably 5 to 10. The multimer may include one or more segments of double-stranded sequences.

Multimeric nucleic acid molecules are useful in amplifying the signal that results from hybridization of one the first sequence of the multimeric molecule to a target sequence. The amplification is theoretically proportional to the number of iterations of the second segment.

Without being held to theory, forked structures of greater than about eight branches exhibited steric hindrance which inhibited binding of labeled probes to the multimer. On the other hand, comb structures exhibit little or no steric problems and are thus a preferred type of branched multimer. For a description of branched nucleic acid multimers of both the fork and comb types, as well as methods of use and synthesis, see, e.g., U.S. Pat. Nos. 5,124,246 (fork-type structures); 5,710,264 (synthesis of comb structures); and 5,849,481.

Use of Polynucleotide Probes in Mapping, and in Tissue Profiling. Polynucleotide probes, generally comprising at least 12 contiguous nt of a polynucleotide as shown in the Sequence Listing, are used for a variety of purposes, such as chromosome mapping of the polynucleotide and detection of transcription levels. Additional disclosure about preferred regions of the disclosed polynucleotide sequences is found in the Examples. A probe that hybridizes specifically to a polynucleotide disclosed herein should provide a detection signal at least 5-, 10-, or 20-fold higher than the background hybridization provided with other unrelated sequences.

Detection of Expression Levels. Nucleotide probes are used to detect expression of a gene corresponding to the provided polynucleotide. In Northern blots, mRNA is separated electrophoretically and contacted with a probe. A probe is detected as hybridizing to an mRNA species of a particular size. The amount of hybridization is quantitated to determine relative amounts of expression, for example under a particular condition. Probes are used for in situ hybridization to cells to detect expression. Probes can also be used in vivo for diagnostic detection of hybridizing sequences. Probes are typically labeled with a radioactive isotope. Other types of detectable labels can be used such as chromophores, fluors, and enzymes. Other examples of nucleotide hybridization assays are described in WO92/02526 and USPN 5,124,246.

Alternatively, the Polymerase Chain Reaction (PCR) is another means for detecting small amounts of target nucleic acids (see, e.g., Mullis et al., *Meth. Enzymol.* (1987) 155:335; USPN 4,683,195; and USPN 4,683,202). Two primer polynucleotides nucleotides that hybridize with the target nucleic acids are used to prime the reaction. The primers can be composed of sequence within or 3' and 5' to the polynucleotides of the Sequence Listing. Alternatively, if the primers are 3' and 5' to these polynucleotides, they need not hybridize to them or the complements. After amplification of the target with a thermostable polymerase, the amplified target nucleic acids can be detected by methods known in the art, e.g., Southern blot. mRNA or cDNA can also be detected by traditional blotting techniques (e.g., Southern blot, Northern blot, etc.) described in Sambrook et al., "Molecular Cloning: A Laboratory Manual" (New York, Cold Spring Harbor Laboratory, 1989) (e.g., without PCR amplification). In general, mRNA or cDNA generated from mRNA using a polymerase enzyme can be purified and separated using gel electrophoresis, and transferred to a solid support, such as nitrocellulose. The solid support is exposed to a labeled probe, washed to remove any unhybridized probe, and duplexes containing the labeled probe are detected.

Mapping. Polynucleotides of the present invention can be used to identify a chromosome on which the corresponding gene resides. Such mapping can be useful in identifying the function of the polynucleotide-related gene by its proximity to other genes with known function. Function can also be assigned to the polynucleotide-related gene when particular syndromes or diseases map to the same chromosome. For example, use of polynucleotide probes in identification and quantification of nucleic acid sequence aberrations is described in USPN 5,783,387. An exemplary mapping method is fluorescence in situ hybridization (FISH), which facilitates comparative genomic hybridization to allow total genome assessment of changes in relative copy number of DNA sequences (see, e.g., Valdes et al., *Methods in Molecular Biology* (1997) 68:1). Polynucleotides can also be mapped to particular chromosomes using, for example, radiation hybrids or chromosome-specific hybrid panels. See Leach et al., *Advances in Genetics*, (1995) 33:63-99; Walter et al., *Nature Genetics* (1994) 7:22; Walter and Goodfellow, *Trends in Genetics* (1992) 9:352. Panels for radiation hybrid mapping are available from Research Genetics, Inc., Huntsville, Alabama, USA. Databases for markers using various panels are available via the world wide web at sites supported by the Stanford Human Genome Center (Stanford University) and the Whitehead Institute for Biomedical Research/MIT Center for Genome Research. The statistical program RHMAP can be used to construct a map based on the data from radiation hybridization with a measure of the relative likelihood of one order versus another. RHMAP is available via the world wide web at a site supported by the University of Michigan. In addition, commercial programs are available for identifying regions of chromosomes commonly associated with disease, such as cancer.

Tissue Typing or Profiling. Expression of specific mRNA corresponding to the provided polynucleotides can vary in different cell types and can be tissue-specific. This variation of mRNA

levels in different cell types can be exploited with nucleic acid probe assays to determine tissue types. For example, PCR, branched DNA probe assays, or blotting techniques utilizing nucleic acid probes substantially identical or complementary to polynucleotides listed in the Sequence Listing can determine the presence or absence of the corresponding cDNA or mRNA.

5 Tissue typing can be used to identify the developmental organ or tissue source of a metastatic lesion by identifying the expression of a particular marker of that organ or tissue. If a polynucleotide is expressed only in a specific tissue type, and a metastatic lesion is found to express that polynucleotide, then the developmental source of the lesion has been identified. Expression of a particular polynucleotide can be assayed by detection of either the corresponding mRNA or the  
10 protein product. As would be readily apparent to any forensic scientist, the sequences disclosed herein are useful in differentiating human tissue from non-human tissue. In particular, these sequences are useful to differentiate human tissue from bird, reptile, and amphibian tissue, for example.

Use of Polymorphisms. A polynucleotide of the invention can be used in forensics, genetic analysis, mapping, and diagnostic applications where the corresponding region of a gene is  
15 polymorphic in the human population. Any means for detecting a polymorphism in a gene can be used, including, but not limited to electrophoresis of protein polymorphic variants, differential sensitivity to restriction enzyme cleavage, and hybridization to allele-specific probes.

Antibody Production. The present invention further provides antibodies, which may be isolated antibodies, that are specific for a polypeptide encoded by a polynucleotide described herein  
20 (e.g., a polypeptide encoded by a sequence corresponding to SEQ ID NOS:1-1485, a polypeptide comprising an amino acid sequence of SEQ ID NOS:1486-1542). Antibodies can be provided in a composition comprising the antibody and a buffer and/or a pharmaceutically acceptable excipient. Antibodies specific for a polypeptide associated with prostate cancer are useful in a variety of diagnostic and therapeutic methods, as discussed in detail herein.

25 Expression products of a polynucleotide of the invention, as well as the corresponding mRNA, cDNA, or complete gene, can be prepared and used for raising antibodies for experimental, diagnostic, and therapeutic purposes. For polynucleotides to which a corresponding gene has not been assigned, this provides an additional method of identifying the corresponding gene. The polynucleotide or related cDNA is expressed as described above, and antibodies are prepared. These  
30 antibodies are specific to an epitope on the polypeptide encoded by the polynucleotide, and can precipitate or bind to the corresponding native protein in a cell or tissue preparation or in a cell-free extract of an in vitro expression system.

Methods for production of antibodies that specifically bind a selected antigen are well known in the art. Immunogens for raising antibodies can be prepared by mixing a polypeptide encoded by a  
35 polynucleotide of the invention with an adjuvant, and/or by making fusion proteins with larger immunogenic proteins. Polypeptides can also be covalently linked to other larger immunogenic

proteins, such as keyhole limpet hemocyanin. Immunogens are typically administered intradermally, subcutaneously, or intramuscularly to experimental animals such as rabbits, sheep, and mice, to generate antibodies. Monoclonal antibodies can be generated by isolating spleen cells and fusing myeloma cells to form hybridomas. Alternatively, the selected polynucleotide is administered directly, such as by intramuscular injection, and expressed in vivo. The expressed protein generates a variety of protein-specific immune responses, including production of antibodies, comparable to administration of the protein.

Preparations of polyclonal and monoclonal antibodies specific for polypeptides encoded by a selected polynucleotide are made using standard methods known in the art. The antibodies specifically bind to epitopes present in the polypeptides encoded by polynucleotides disclosed in the Sequence Listing. Typically, at least 6, 8, 10, or 12 contiguous amino acids are required to form an epitope. Epitopes that involve non-contiguous amino acids may require a longer polypeptide, e.g., at least 15, 25, or 50 amino acids. Antibodies that specifically bind to human polypeptides encoded by the provided polypeptides should provide a detection signal at least 5-, 10-, or 20-fold higher than a detection signal provided with other proteins when used in Western blots or other immunochemical assays. Preferably, antibodies that specifically bind polypeptides contemplated by the invention do not bind to other proteins in immunochemical assays at detectable levels and can immunoprecipitate the specific polypeptide from solution.

The invention also contemplates naturally occurring antibodies specific for a polypeptide of the invention. For example, serum antibodies to a polypeptide of the invention in a human population can be purified by methods well known in the art, e.g., by passing antiserum over a column to which the corresponding selected polypeptide or fusion protein is bound. The bound antibodies can then be eluted from the column, for example, using a buffer with a high salt concentration.

In addition to the antibodies discussed above, the invention also contemplates genetically engineered antibodies (e.g., chimeric antibodies, humanized antibodies, human antibodies produced by a transgenic animal (e.g., a transgenic mouse such as the Xenomous™), antibody derivatives (e.g., single chain antibodies, antibody fragments (e.g., Fab, etc.)), according to methods well known in the art.

The invention also contemplates other molecules that can specifically bind a polynucleotide or polypeptide of the invention. Examples of such molecules include, but are not necessarily limited to, single-chain binding proteins (e.g., mono- and multi-valent single chain antigen binding proteins (see, e.g., U.S. Patent Nos. 4,704,692; 4,946,778; 4,946,778; 6,027,725; 6,121,424)), oligonucleotide-based synthetic antibodies (e.g., oligobodies (see, e.g., Radrizzani et al., *Medicina* (B Aires) (1999) 59:753-8; Radrizzani et al., *Medicina* (B Aires) (2000) 60(Suppl 2):55-60)), aptamers (see, e.g., Gening et al., *Biotechniques* (2001) 3:828, 830, 832, 834; Cox and Ellington, *Bioorg. Med. Chem.* (2001) 9:2525-31), and the like.

### Polynucleotides or Arrays for Diagnostics.

Polynucleotide arrays provide a high throughput technique that can assay a large number of polynucleotides in a sample. This technology can be used as a diagnostic and as tool to test for differential expression expression, e.g., to determine function of an encoded protein. A variety of methods of producing arrays, as well as variations of these methods, are known in the art and contemplated for use in the invention. For example, arrays can be created by spotting polynucleotide probes onto a substrate (e.g., glass, nitrocellulose, etc.) in a two-dimensional matrix or array having bound probes. The probes can be bound to the substrate by either covalent bonds or by non-specific interactions, such as hydrophobic interactions. Samples of polynucleotides can be detectably labeled (e.g., using radioactive or fluorescent labels) and then hybridized to the probes. Double stranded polynucleotides, comprising the labeled sample polynucleotides bound to probe polynucleotides, can be detected once the unbound portion of the sample is washed away. Alternatively, the polynucleotides of the test sample can be immobilized on the array, and the probes detectably labeled.

Techniques for constructing arrays and methods of using these arrays are described in, for example, Schena et al. (1996) Proc Natl Acad Sci U S A. 93(20):10614-9; Schena et al. (1995) Science 270(5235):467-70; Shalon et al. (1996) Genome Res. 6(7):639-45, USPN 5,807,522, EP 799 897; WO 97/29212; WO 97/27317; EP 785 280; WO 97/02357; USPN 5,593,839; USPN 5,578,832; EP 728 520; USPN 5,599,695; EP 721 016; USPN 5,556,752; WO 95/22058; and USPN 5,631,734.

Arrays can be used to, for example, examine differential expression of genes and can be used to determine gene function. For example, arrays can be used to detect differential expression of a gene corresponding to a polynucleotide of the invention, where expression is compared between a test cell and control cell (e.g., cancer cells and normal cells). For example, high expression of a particular message in a cancer cell, which is not observed in a corresponding normal cell, can indicate a cancer specific gene product. Exemplary uses of arrays are further described in, for example, Pappalardo et al., Sem. Radiation Oncol. (1998) 8:217; and Ramsay Nature Biotechnol. (1998) 16:40. Furthermore, many variations on methods of detection using arrays are well within the skill in the art and within the scope of the present invention. For example, rather than immobilizing the probe to a solid support, the test sample can be immobilized on a solid support which is then contacted with the probe.

### Differential Expression in Diagnosis

The polynucleotides of the invention can also be used to detect differences in expression levels between two cells, e.g., as a method to identify abnormal or diseased tissue in a human. For polynucleotides corresponding to profiles of protein families, the choice of tissue can be selected according to the putative biological function. In general, the expression of a gene corresponding to a specific polynucleotide is compared between a first tissue that is suspected of being diseased and a second, normal tissue of the human. The tissue suspected of being abnormal or diseased can be derived from a different tissue type of the human, but preferably it is derived from the same tissue



type; for example, an intestinal polyp or other abnormal growth should be compared with normal intestinal tissue. The normal tissue can be the same tissue as that of the test sample, or any normal tissue of the patient, especially those that express the polynucleotide-related gene of interest (e.g., brain, thymus, testis, heart, prostate, placenta, spleen, small intestine, skeletal muscle, pancreas, and the mucosal lining of the colon). A difference between the polynucleotide-related gene, mRNA, or protein in the two tissues which are compared, for example, in molecular weight, amino acid or nucleotide sequence, or relative abundance, indicates a change in the gene, or a gene which regulates it, in the tissue of the human that was suspected of being diseased. Examples of detection of differential expression and its use in diagnosis of cancer are described in USPNs 5,688,641 and 5,677,125.

A genetic predisposition to disease in a human can also be detected by comparing expression levels of an mRNA or protein corresponding to a polynucleotide of the invention in a fetal tissue with levels associated in normal fetal tissue. Fetal tissues that are used for this purpose include, but are not limited to, amniotic fluid, chorionic villi, blood, and the blastomere of an in vitro-fertilized embryo.

The comparable normal polynucleotide-related gene is obtained from any tissue. The mRNA or protein is obtained from a normal tissue of a human in which the polynucleotide-related gene is expressed. Differences such as alterations in the nucleotide sequence or size of the same product of the fetal polynucleotide-related gene or mRNA, or alterations in the molecular weight, amino acid sequence, or relative abundance of fetal protein, can indicate a germline mutation in the polynucleotide-related gene of the fetus, which indicates a genetic predisposition to disease. In general, diagnostic, prognostic, and other methods of the invention based on differential expression involve detection of a level or amount of a gene product, particularly a differentially expressed gene product, in a test sample obtained from a patient suspected of having or being susceptible to a disease (e.g., breast cancer, prostate cancer, lung cancer, colon cancer and/or metastatic forms thereof), and comparing the detected levels to those levels found in normal cells (e.g., cells substantially unaffected by cancer) and/or other control cells (e.g., to differentiate a cancerous cell from a cell affected by dysplasia). Furthermore, the severity of the disease can be assessed by comparing the detected levels of a differentially expressed gene product with those levels detected in samples representing the levels of differentially expressed gene product associated with varying degrees of severity of disease. It should be noted that use of the term "diagnostic" herein is not necessarily meant to exclude "prognostic" or "prognosis," but rather is used as a matter of convenience.

The term "differentially expressed gene" is generally intended to encompass a polynucleotide that can, for example, include an open reading frame encoding a gene product (e.g., a polypeptide), and/or introns of such genes and adjacent 5' and 3' non-coding nucleotide sequences involved in the regulation of expression, up to about 20 kb beyond the coding region, but possibly further in either direction. The gene can be introduced into an appropriate vector for extrachromosomal maintenance

or for integration into a host genome. In general, a difference in expression level associated with a decrease in expression level of at least about 25%, usually at least about 50% to 75%, more usually at least about 90% or more is indicative of a differentially expressed gene of interest, i.e., a gene that is underexpressed or down-regulated in the test sample relative to a control sample. Furthermore, a difference in expression level associated with an increase in expression of at least about 25%, usually at least about 50% to 75%, more usually at least about 90% and can be at least about 1½-fold, usually at least about 2-fold to about 10-fold, and can be about 100-fold to about 1,000-fold increase relative to a control sample is indicative of a differentially expressed gene of interest, i.e., an overexpressed or up-regulated gene.

"Differentially expressed polynucleotide" as used herein means a nucleic acid molecule (RNA or DNA) comprising a sequence that represents a differentially expressed gene, e.g., the differentially expressed polynucleotide comprises a sequence (e.g., an open reading frame encoding a gene product) that uniquely identifies a differentially expressed gene so that detection of the differentially expressed polynucleotide in a sample is correlated with the presence of a differentially expressed gene in a sample. "Differentially expressed polynucleotide" is also meant to encompass fragments of the disclosed polynucleotides, e.g., fragments retaining biological activity, as well as nucleic acids homologous, substantially similar, or substantially identical (e.g., having about 90% sequence identity) to the disclosed polynucleotides.

Methods of the subject invention useful in diagnosis or prognosis typically involve comparison of the abundance of a selected differentially expressed gene product in a sample of interest with that of a control to determine any relative differences in the expression of the gene product, where the difference can be measured qualitatively and/or quantitatively. Quantitation can be accomplished, for example, by comparing the level of expression product detected in the sample with the amounts of product present in a standard curve. A comparison can be made visually, by using a technique such as densitometry, with or without computerized assistance; by preparing a representative library of cDNA clones of mRNA isolated from a test sample, sequencing the clones in the library to determine that number of cDNA clones corresponding to the same gene product, and analyzing the number of clones corresponding to that same gene product relative to the number of clones of the same gene product in a control sample; or by using an array to detect relative levels of hybridization to a selected sequence or set of sequences, and comparing the hybridization pattern to that of a control. The differences in expression are then correlated with the presence or absence of an abnormal expression pattern. A variety of different methods for determining the nucleic acid abundance in a sample are known to those of skill in the art (see, e.g., WO 97/27317).

In general, diagnostic assays of the invention involve detection of a gene product of a polynucleotide sequence (e.g., mRNA or polypeptide) that corresponds to a sequence of SEQ ID NOS:1-1485. The patient from whom the sample is obtained can be apparently healthy, susceptible to

disease (e.g., as determined by family history or exposure to certain environmental factors), or can already be identified as having a condition in which altered expression of a gene product of the invention is implicated.

Diagnosis can be determined based on detected gene product expression levels of a gene product encoded by at least one, preferably at least two or more, at least 3 or more, or at least 4 or more of the polynucleotides having a sequence set forth in SEQ ID NOS:1-1485, and can involve detection of expression of genes corresponding to all of SEQ ID NOS:1-1485 and/or additional sequences that can serve as additional diagnostic markers and/or reference sequences. Where the diagnostic method is designed to detect the presence or susceptibility of a patient to cancer, the assay preferably involves detection of a gene product encoded by a gene corresponding to a polynucleotide that is differentially expressed in cancer. Examples of such differentially expressed polynucleotides are described in the Examples below. Given the provided polynucleotides and information regarding their relative expression levels provided herein, assays using such polynucleotides and detection of their expression levels in diagnosis and prognosis will be readily apparent to the ordinarily skilled artisan.

Any of a variety of detectable labels can be used in connection with the various embodiments of the diagnostic methods of the invention. Suitable detectable labels include fluorochromes, (e.g. fluorescein isothiocyanate (FITC), rhodamine, Texas Red, phycoerythrin, allophycocyanin, 6-carboxyfluorescein (6-FAM), 2',7'-dimethoxy-4',5'-dichloro-6-carboxyfluorescein, 6-carboxy-X-rhodamine (ROX), 6-carboxy-2',4',7',4,7-hexachlorofluorescein (HEX), 5-carboxyfluorescein (5-FAM) or N,N,N',N'-tetramethyl-6-carboxyrhodamine (TAMRA)), radioactive labels, (e.g. 32P, 35S, 3H, etc.), and the like. The detectable label can involve a two stage systems (e.g., biotin-avidin, hapten-anti-hapten antibody, etc.).

Reagents specific for the polynucleotides and polypeptides of the invention, such as antibodies and nucleotide probes, can be supplied in a kit for detecting the presence of an expression product in a biological sample. The kit can also contain buffers or labeling components, as well as instructions for using the reagents to detect and quantify expression products in the biological sample. Exemplary embodiments of the diagnostic methods of the invention are described below in more detail.

Polypeptide detection in diagnosis. In one embodiment, the test sample is assayed for the level of a differentially expressed polypeptide, such as a polypeptide of a gene corresponding to SEQ ID NOS:1-1485 and/or a polypeptide comprising a sequence of SEQ ID NO:1486-1542. Diagnosis can be accomplished using any of a number of methods to determine the absence or presence or altered amounts of the differentially expressed polypeptide in the test sample. For example, detection can utilize staining of cells or histological sections with labeled antibodies, performed in accordance with conventional methods. Cells can be permeabilized to stain cytoplasmic molecules. In general,

antibodies that specifically bind a differentially expressed polypeptide of the invention are added to a sample, and incubated for a period of time sufficient to allow binding to the epitope, usually at least about 10 minutes. The antibody can be detectably labeled for direct detection (e.g., using radioisotopes, enzymes, fluorescers, chemilumescers, and the like), or can be used in conjunction with a second stage antibody or reagent to detect binding (e.g., biotin with horseradish peroxidase-conjugated avidin, a secondary antibody conjugated to a fluorescent compound, e.g. fluorescein, rhodamine, Texas red, etc.). The absence or presence of antibody binding can be determined by various methods, including flow cytometry of dissociated cells, microscopy, radiography, scintillation counting, etc. Any suitable alternative methods of qualitative or quantitative detection of levels or amounts of differentially expressed polypeptide can be used, for example, ELISA, western blot, immunoprecipitation, radioimmunoassay, etc.

mRNA detection. The diagnostic methods of the invention can also or alternatively involve detection of mRNA encoded by a gene corresponding to a differentially expressed polynucleotide of the invention. Any suitable qualitative or quantitative methods known in the art for detecting specific mRNAs can be used. mRNA can be detected by, for example, in situ hybridization in tissue sections, by reverse transcriptase-PCR, or in Northern blots containing poly A+ mRNA. One of skill in the art can readily use these methods to determine differences in the size or amount of mRNA transcripts between two samples. mRNA expression levels in a sample can also be determined by generation of a library of expressed sequence tags (ESTs) from the sample, where the EST library is representative of sequences present in the sample (Adams et al., (1991) *Science* 252:1651). Enumeration of the relative representation of ESTs within the library can be used to approximate the relative representation of the gene transcript within the starting sample. The results of EST analysis of a test sample can then be compared to EST analysis of a reference sample to determine the relative expression levels of a selected polynucleotide, particularly a polynucleotide corresponding to one or more of the differentially expressed genes described herein. Alternatively, gene expression in a test sample can be performed using serial analysis of gene expression (SAGE) methodology (e.g., Velculescu et al., *Science* (1995) 270:484) or differential display (DD) methodology (see, e.g., USPN 5,776,683 and USPN 5,807,680).

Alternatively, gene expression can be analyzed using hybridization analysis. Oligonucleotides or cDNA can be used to selectively identify or capture DNA or RNA of specific sequence composition, and the amount of RNA or cDNA hybridized to a known capture sequence determined qualitatively or quantitatively, to provide information about the relative representation of a particular message within the pool of cellular messages in a sample. Hybridization analysis can be designed to allow for concurrent screening of the relative expression of hundreds to thousands of genes by using, for example, array-based technologies having high density formats, including filters, microscope slides, or microchips, or solution-based technologies that use spectroscopic analysis (e.g., mass

spectrometry). One exemplary use of arrays in the diagnostic methods of the invention is described below in more detail.

Use of a single gene in diagnostic applications. The diagnostic methods of the invention can focus on the expression of a single differentially expressed gene. For example, the diagnostic method can involve detecting a differentially expressed gene, or a polymorphism of such a gene (e.g., a polymorphism in a coding region or control region), that is associated with disease. Disease-associated polymorphisms can include deletion or truncation of the gene, mutations that alter expression level and/or affect activity of the encoded protein, etc.

A number of methods are available for analyzing nucleic acids for the presence of a specific sequence, e.g. a disease associated polymorphism. Where large amounts of DNA are available, genomic DNA is used directly. Alternatively, the region of interest is cloned into a suitable vector and grown in sufficient quantity for analysis. Cells that express a differentially expressed gene can be used as a source of mRNA, which can be assayed directly or reverse transcribed into cDNA for analysis. The nucleic acid can be amplified by conventional techniques, such as the polymerase chain reaction (PCR), to provide sufficient amounts for analysis, and a detectable label can be included in the amplification reaction (e.g., using a detectably labeled primer or detectably labeled oligonucleotides) to facilitate detection. Alternatively, various methods are also known in the art that utilize oligonucleotide ligation as a means of detecting polymorphisms, see, e.g., Riley et al., Nucl. Acids Res. (1990) 18:2887; and Delahunty et al., Am. J. Hum. Genet. (1996) 58:1239.

The amplified or cloned sample nucleic acid can be analyzed by one of a number of methods known in the art. The nucleic acid can be sequenced by dideoxy or other methods, and the sequence of bases compared to a selected sequence, e.g., to a wild-type sequence. Hybridization with the polymorphic or variant sequence can also be used to determine its presence in a sample (e.g., by Southern blot, dot blot, etc.). The hybridization pattern of a polymorphic or variant sequence and a control sequence to an array of oligonucleotide probes immobilized on a solid support, as described in US 5,445,934, or in WO 95/35505, can also be used as a means of identifying polymorphic or variant sequences associated with disease. Single strand conformational polymorphism (SSCP) analysis, denaturing gradient gel electrophoresis (DGGE), and heteroduplex analysis in gel matrices are used to detect conformational changes created by DNA sequence variation as alterations in electrophoretic mobility. Alternatively, where a polymorphism creates or destroys a recognition site for a restriction endonuclease, the sample is digested with that endonuclease, and the products size fractionated to determine whether the fragment was digested. Fractionation is performed by gel or capillary electrophoresis, particularly acrylamide or agarose gels.

Screening for mutations in a gene can be based on the functional or antigenic characteristics of the protein. Protein truncation assays are useful in detecting deletions that can affect the biological activity of the protein. Various immunoassays designed to detect polymorphisms in proteins can be

used in screening. Where many diverse genetic mutations lead to a particular disease phenotype, functional protein assays have proven to be effective screening tools. The activity of the encoded protein can be determined by comparison with the wild-type protein.

Diagnosis, Prognosis, Assessment of Therapy (Therapeutics), and Management of Cancer

5 The polynucleotides of the invention, as well as their gene products, are of particular interest as genetic or biochemical markers (e.g., in blood or tissues) that will detect the earliest changes along the carcinogenesis pathway and/or to monitor the efficacy of various therapies and preventive interventions. For example, the level of expression of certain polynucleotides can be indicative of a poorer prognosis, and therefore warrant more aggressive chemo- or radio-therapy for a patient or vice versa. The correlation of novel surrogate tumor specific features with response to treatment and outcome in patients can define prognostic indicators that allow the design of tailored therapy based on the molecular profile of the tumor. These therapies include antibody targeting, antagonists (e.g., small molecules), and gene therapy. Determining expression of certain polynucleotides and comparison of a patient's profile with known expression in normal tissue and variants of the disease allows a determination of the best possible treatment for a patient, both in terms of specificity of treatment and in terms of comfort level of the patient. Surrogate tumor markers, such as polynucleotide expression, can also be used to better classify, and thus diagnose and treat, different forms and disease states of cancer. Two classifications widely used in oncology that can benefit from identification of the expression levels of the genes corresponding to the polynucleotides of the invention are staging of the cancerous disorder, and grading the nature of the cancerous tissue.

20 The polynucleotides that correspond to differentially expressed genes, as well as their encoded gene products, can be useful to monitor patients having or susceptible to cancer to detect potentially malignant events at a molecular level before they are detectable at a gross morphological level. In addition, the polynucleotides of the invention, as well as the genes corresponding to such polynucleotides, can be useful as therapeutics, e.g., to assess the effectiveness of therapy by using the polynucleotides or their encoded gene products, to assess, for example, tumor burden in the patient before, during, and after therapy.

25 Furthermore, a polynucleotide identified as corresponding to a gene that is differentially expressed in, and thus is important for, one type of cancer can also have implications for development or risk of development of other types of cancer, e.g., where a polynucleotide represents a gene differentially expressed across various cancer types. Thus, for example, expression of a polynucleotide corresponding to a gene that has clinical implications for metastatic colon cancer can also have clinical implications for stomach cancer or endometrial cancer.

30 Staging. Staging is a process used by physicians to describe how advanced the cancerous state is in a patient. Staging assists the physician in determining a prognosis, planning treatment and evaluating the results of such treatment. Staging systems vary with the types of cancer, but generally

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involve the following "TNM" system: the type of tumor, indicated by T; whether the cancer has metastasized to nearby lymph nodes, indicated by N; and whether the cancer has metastasized to more distant parts of the body, indicated by M. Generally, if a cancer is only detectable in the area of the primary lesion without having spread to any lymph nodes, it is called Stage I or Stage II, depending on the degree of invasiveness as indicated by the tumor grade of the primary lesion. If the primary lesion is of tumor grade I or II and the patient does not have any regional or distant metastasis, the cancer is classified as Stage I. If the primary lesion is of tumor grade III or IV and the patient does not have any regional or distant metastasis, the cancer is classified as Stage II. If the cancer has spread only to the regional lymph nodes, it is classified as Stage III. Cancers that have spread to a distant part of the body, such as liver, bone, brain or other sites, are Stage IV, the most advanced stage.

The polynucleotides of the invention can facilitate fine-tuning of the staging process by identifying markers for the aggressivity of a cancer, e.g., the metastatic potential, as well as the presence in different areas of the body. Thus, a Stage II cancer with a polynucleotide signifying a high metastatic potential cancer can be used to change a borderline Stage II tumor to a Stage III tumor, justifying more aggressive therapy. Conversely, the presence of a polynucleotide signifying a lower metastatic potential allows more conservative staging of a tumor.

Grading of cancers. Grade is a term used to describe how closely a tumor resembles normal tissue of its same type. The microscopic appearance of a tumor is used to identify tumor grade based on parameters such as cell morphology, cellular organization, and other markers of differentiation. As a general rule, the grade of a tumor corresponds to its rate of growth or aggressiveness, with undifferentiated or high-grade tumors being more aggressive than well-differentiated or low-grade tumors. The following guidelines are generally used for grading tumors: 1) GX Grade cannot be assessed; 2) G1 Well differentiated; 3) G2 Moderately well differentiated; 4) G3 Poorly differentiated; 5) G4 Undifferentiated. The polynucleotides of the invention can be especially valuable in determining the grade of the tumor, as they not only can aid in determining the differentiation status of the cells of a tumor, they can also identify factors other than differentiation that are valuable in determining the aggressiveness of a tumor, such as metastatic potential.

For prostate cancer, the Gleason Grading/Scoring system is most commonly used. A prostate biopsy tissue sample is examined under a microscope and a grade is assigned to the tissue based on: 1) the appearance of the cells, and 2) the arrangement of the cells. Each parameter is assessed on a scale of one (cells are almost normal) to five (abnormal), and the individual Gleason Grades are presented separated by a "+" sign. Alternatively, the two grades are combined to give a Gleason Score of 2-10. Thus, for a tissue sample that received a grade of 3 for each parameter, the Gleason Grade would be 3+3 and the Gleason Score would be 6. A lower Gleason Score indicates a well-differentiated tumor, while a higher Gleason Score indicates a poorly differentiated cancer that is more likely to spread. The majority of biopsies in general are Gleason Scores 5, 6 and 7.

Gleason Score 2, 3, 4	Gleason Score 5, 6, 7	Gleason Score 8, 9, 10
Low-grade tumor	Medium-grade tumor	High-grade tumor
Slow Growth	Unpredictable Growth	Aggressive Growth
Least dangerous.  Cells look most like normal prostate cells and are described as being "well-differentiated".  Tends to be slow growing.	Intermediate cancers may behave like low-grade or high-grade cancers.  The cells' behavior may depend on the volume of the cancer and the PSA level.  This is the most common grade of prostate cancer.	High-grade cancers are usually very aggressive and quick to spread to the tissue surrounding the prostate.  These cancer cells look least like normal prostate cells and are usually described as "poorly-differentiated".

The polynucleotides of the Sequence Listing, and their corresponding genes and gene products, can be especially valuable in determining the grade of the tumor, as they not only can aid in determining the differentiation status of the cells of a tumor, they can also identify factors other than differentiation that are valuable in determining the aggressiveness of a tumor, such as metastatic potential.

Assessment of proliferation of cells in tumor. The differential expression level of the polynucleotides described herein can facilitate assessment of the rate of proliferation of tumor cells, and thus provide an indicator of the aggressiveness of the rate of tumor growth. For example, assessment of the relative expression levels of genes involved in the cell cycle can provide an indication of cellular proliferation, and thus serve as a marker of proliferation.

Detection of colon cancer. The polynucleotides corresponding to genes that exhibit the appropriate expression pattern can be used to detect colon cancer in a subject. Colorectal cancer is one of the most common neoplasms in humans and perhaps the most frequent form of hereditary neoplasia. Prevention and early detection are key factors in controlling and curing colorectal cancer. Colorectal cancer begins as polyps, which are small, benign growths of cells that form on the inner lining of the colon. Over a period of several years, some of these polyps accumulate additional mutations and become cancerous. Multiple familial colorectal cancer disorders have been identified, which are summarized as follows: 1) Familial adenomatous polyposis (FAP); 2) Gardner's syndrome; 3) Hereditary nonpolyposis colon cancer (HNPCC); and 4) Familial colorectal cancer in Ashkenazi Jews. The expression of appropriate polynucleotides of the invention can be used in the diagnosis, prognosis and management of colorectal cancer. Detection of colon cancer can be determined using expression levels of any of these sequences alone or in combination with the levels of expression. Determination of the aggressive nature and/or the metastatic potential of a colon cancer can be determined by comparing levels of one or more polynucleotides of the invention and comparing total levels of another sequence known to vary in cancerous tissue, e.g., expression of p53, DCC ras, or FAP (see, e.g., Fearon ER, et al., Cell (1990) 61(5):759; Hamilton SR et al., Cancer (1993) 72:957;



Bodmer W, et al., Nat Genet. (1994) 4(3):217; Fearon ER, Ann N Y Acad Sci. (1995) 768:101). For example, development of colon cancer can be detected by examining the ratio of any of the polynucleotides of the invention to the levels of oncogenes (e.g., ras) or tumor suppressor genes (e.g., FAP or p53). Thus, expression of specific marker polynucleotides can be used to discriminate between normal and cancerous colon tissue, to discriminate between colon cancers with different cells of origin, to discriminate between colon cancers with different potential metastatic rates, etc. For a review of markers of cancer, see, e.g., Hanahan et al. (2000) Cell 100:57-70.

Detection of prostate cancer. The polynucleotides and their corresponding genes and gene products exhibiting the appropriate differential expression pattern can be used to detect prostate cancer in a subject. Prostate cancer is quite common in humans, with one out of every six men at a lifetime risk for prostate cancer, and can be relatively harmless or extremely aggressive. Some prostate tumors are slow growing, causing few clinical symptoms, while aggressive tumors spread rapidly to the lymph nodes, other organs and especially bone. Over 95% of primary prostate cancers are adenocarcinomas. Signs and symptoms may include: frequent urination, especially at night; inability to urinate; trouble starting or holding back urination; a weak or interrupted urine flow; and frequent pain or stiffness in the lower back, hips or upper thighs.

The prostate is divided into three areas - the peripheral zone, the transition zone, and the central zone - with a layer of tissue surrounding all three. Most prostate tumors form in the peripheral zone; the larger, glandular portion of the organ. Prostate cancer can also form in the tissue of the central zone. Surrounding the prostate is the prostate capsule, a tissue that separates the prostate from the rest of the body. When prostate cancer remains inside the prostate capsule, it is considered localized and treatable with surgery. Once the cancer punctures the capsule and spreads outside, treatment options are more limited. Prevention and early detection are key factors in controlling and curing prostate cancer.

While the Gleason Grade or Score of a prostate cancer can provide information useful in determining the appropriate treatment of a prostate cancer, the majority of prostate cancers are Gleason Scores 5, 6, and 7, which exhibit unpredictable behavior. These cancers may behave like less dangerous low-grade cancers or like extremely dangerous high-grade cancers. As a result, a patient living with a medium-grade prostate cancer is at constant risk of developing high-grade cancer.

The expression of appropriate polynucleotides can be used in the diagnosis, prognosis and management of prostate cancer. Detection of prostate cancer can be determined using expression levels of any of these sequences alone or in combination with the levels of expression of any other nucleotide sequences. Determination of the aggressive nature and/or the metastatic potential of a prostate cancer can be determined by comparing levels of one or more gene products of the genes corresponding to the polynucleotides described herein, and comparing total levels of another sequence known to vary in cancerous tissue, e.g., expression of p53, DCC, ras, FAP (see, e.g., Fearon ER, et

*al., Cell* (1990) 61(5):759; Hamilton SR *et al., Cancer* (1993) 72:957; Bodmer W, *et al., Nat Genet.* (1994) 4(3):217; Fearon ER, *Ann NY Acad Sci.* (1995) 768:101).

For example, development of prostate cancer can be detected by examining the level of expression of a gene corresponding to a polynucleotides described herein to the levels of oncogenes (e.g. ras) or tumor suppressor genes (e.g. FAP or p53). Thus expression of specific marker polynucleotides can be used to discriminate between normal and cancerous prostate tissue, to discriminate between prostate cancers with different cells of origin, to discriminate between prostate cancers with different potential metastatic rates, etc. For a review of markers of cancer, see, e.g., Hanahan *et al.* (2000) *Cell* 100:57-70.

In addition, many of the signs and symptoms of prostate cancer can be caused by a variety of other non-cancerous conditions. For example, one common cause of many of these signs and symptoms is a condition called benign prostatic hypertrophy, or BPH. In BPH, the prostate gets bigger and may block the flow of urine or interfere with sexual function. The methods and compositions of the invention can be used to distinguish between prostate cancer and such non-cancerous conditions. The methods of the invention can be used in conjunction with conventional methods of diagnosis, e.g., digital rectal exam and/or detection of the level of prostate specific antigen (PSA), a substance produced and secreted by the prostate, and/or prostatic acid phosphatase (PAP).

Detection of breast cancer. The majority of breast cancers are adenocarcinoma subtypes, which can be summarized as follows: 1) ductal carcinoma in situ (DCIS), including comedocarcinoma; 2) infiltrating (or invasive) ductal carcinoma (IDC); 3) lobular carcinoma in situ (LCIS); 4) infiltrating (or invasive) lobular carcinoma (ILC); 5) inflammatory breast cancer; 6) medullary carcinoma; 7) mucinous carcinoma; 8) Paget's disease of the nipple; 9) Phyllodes tumor; and 10) tubular carcinoma.

The expression of polynucleotides of the invention can be used in the diagnosis and management of breast cancer, as well as to distinguish between types of breast cancer. Detection of breast cancer can be determined using expression levels of any of the appropriate polynucleotides of the invention, either alone or in combination. Determination of the aggressive nature and/or the metastatic potential of a breast cancer can also be determined by comparing levels of one or more polynucleotides of the invention and comparing levels of another sequence known to vary in cancerous tissue, e.g., ER expression. In addition, development of breast cancer can be detected by examining the ratio of expression of a differentially expressed polynucleotide to the levels of steroid hormones (e.g., testosterone or estrogen) or to other hormones (e.g., growth hormone, insulin). Thus, expression of specific marker polynucleotides can be used to discriminate between normal and cancerous breast tissue, to discriminate between breast cancers with different cells of origin, to discriminate between breast cancers with different potential metastatic rates, etc.

Detection of lung cancer. The polynucleotides of the invention can be used to detect lung cancer in a subject. Although there are more than a dozen different kinds of lung cancer, the two main types of lung cancer are small cell and nonsmall cell, which encompass about 90% of all lung cancer cases. Small cell carcinoma (also called oat cell carcinoma) usually starts in one of the larger bronchial tubes, grows fairly rapidly, and is likely to be large by the time of diagnosis. Nonsmall cell lung cancer (NSCLC) is made up of three general subtypes of lung cancer. Epidermoid carcinoma (also called squamous cell carcinoma) usually starts in one of the larger bronchial tubes and grows relatively slowly. The size of these tumors can range from very small to quite large. Adenocarcinoma starts growing near the outside surface of the lung and can vary in both size and growth rate. Some slowly growing adenocarcinomas are described as alveolar cell cancer. Large cell carcinoma starts near the surface of the lung, grows rapidly, and the growth is usually fairly large when diagnosed. Other less common forms of lung cancer are carcinoid, cylindroma, mucoepidermoid, and malignant mesothelioma.

The polynucleotides of the invention, e.g., polynucleotides differentially expressed in normal cells versus cancerous lung cells (e.g., tumor cells of high or low metastatic potential) or between types of cancerous lung cells (e.g., high metastatic versus low metastatic), can be used to distinguish types of lung cancer as well as identifying traits specific to a certain patient's cancer and selecting an appropriate therapy. For example, if the patient's biopsy expresses a polynucleotide that is associated with a low metastatic potential, it may justify leaving a larger portion of the patient's lung in surgery to remove the lesion. Alternatively, a smaller lesion with expression of a polynucleotide that is associated with high metastatic potential may justify a more radical removal of lung tissue and/or the surrounding lymph nodes, even if no metastasis can be identified through pathological examination.

#### Tumor classification and patient stratification

The invention further provides for methods of classifying tumors, and thus grouping or "stratifying" patients, according to the expression profile of selected differentially expressed genes in a tumor. Differentially expressed genes can be analyzed for correlation with other differentially expressed genes in a single tumor type (e.g., a prostate tumor) or between tumor types (e.g., between prostate and colon tumors). Genes that demonstrate consistent correlation in expression profile in a given cancer cell type (e.g., in a prostate cancer cell or type of prostate cancer) can be grouped together, e.g., when one gene is overexpressed in a tumor, a second gene is also usually overexpressed. Tumors can then be classified according to the expression profile of one or more genes selected from one or more groups.

The tumor of each patient in a pool of potential patients can be classified as described above. Patients having similarly classified tumors can then be selected for participation in an investigative or clinical trial of a cancer therapeutic where a homogeneous population is desired. The tumor

classification of a patient can also be used in assessing the efficacy of a cancer therapeutic in a heterogeneous patient population. In addition, therapy for a patient having a tumor of a given expression profile can then be selected accordingly.

#### Treatment of cancer

5 The invention further provides methods for reducing growth of cancer cells. In general, the methods comprise contacting a cancer cell with a substance that modulates (1) expression of a polynucleotide corresponding to a gene that is differentially expressed in cancer; or (2) a level of and/or an activity of a cancer-associated polypeptide. In general, the methods provide for decreasing the expression of a gene that is differentially expressed in a cancer cell (e.g., overexpressed) or  
10 decreasing the level of and/or decreasing an activity of a cancer-associated polypeptide. The methods also provide for increasing expression of a gene that is underexpressed in a cancer cell or increasing the level of and/or increasing an activity of a cancer-associated polypeptide.

"Reducing growth of cancer cells" includes, but is not limited to, reducing proliferation of cancer cells (e.g., prostate, colon, lung, breast, etc. cancer cells), and reducing the incidence of a non-cancerous cell becoming a cancerous cell. Whether a reduction in cancer cell growth has been  
15 achieved can be readily determined using any known assay, including, but not limited to, [<sup>3</sup>H]-thymidine incorporation; counting cell number over a period of time; detecting and/or measuring a marker associated with the cancer type (e.g., CEA, CA19-9, LASA, PSA, PAP, CA15-3, CA27-29, NSE, LDH, etc.).

20 The present invention provides methods for treating cancer, generally comprising administering to an individual in need thereof a substance that reduces cancer cell growth, in an amount sufficient to reduce cancer cell growth and treat the cancer. Whether a substance, or a specific amount of the substance, is effective in treating cancer can be assessed using any of a variety of known diagnostic assays for the particular type of cancer being treated. The substance can be  
25 administered systemically or locally. Thus, in some embodiments, the substance is administered locally, and cancer growth is decreased at the site of administration. Local administration may be useful in treating, e.g., a solid tumor.

A substance that reduces cancer cell growth can be targeted to a cancer cell. Thus, in some embodiments, the invention provides a method of delivering a drug to a cancer cell, comprising  
30 administering a drug-antibody complex to a subject, wherein the antibody is specific for a particular cancer-associated polypeptide, and the drug is one that reduces cancer cell growth, a variety of which are known in the art. Targeting can be accomplished by coupling (e.g., linking, directly or via a linker molecule, either covalently or non-covalently, so as to form a drug-antibody complex) a drug to an antibody specific for a particular cancer-associated polypeptide. Methods of coupling a drug to an  
35 antibody are well known in the art and need not be elaborated upon herein.

In another embodiment, differentially expressed gene products (*e.g.*, polypeptides or polynucleotides encoding such polypeptides) may be effectively used in treatment through vaccination. The growth of cancer cells is naturally limited in part due to immune surveillance. Stimulation of the immune system using a particular tumor-specific antigen enhances the effect  
5 towards the tumor expressing the antigen. An active vaccine comprising a polypeptide encoded by the cDNA of this invention would be appropriately administered to subjects having overabundance of the corresponding RNA, or those predisposed for developing cancer cells with overabundance of the same RNA. Polypeptide antigens are typically combined with an adjuvant as part of a vaccine composition. The vaccine is preferably administered first as a priming dose, and then again as a boosting dose,  
10 usually at least four weeks later. Further boosting doses may be given to enhance the effect. The dose and its timing are usually determined by the person responsible for the treatment.

The invention also encompasses the selection of a therapeutic regimen based upon the expression profile of differentially expressed genes in the patient's tumor. For example, a tumor can be analyzed for its expression profile of the genes corresponding to SEQ ID NOS:1-1542 as described  
15 herein, *e.g.*, the tumor is analyzed to determine which genes are expressed at elevated levels or at decreased levels relative to normal cells of the same tissue type. The expression patterns of the tumor are then compared to the expression patterns of tumors that respond to a selected therapy. Where the expression profiles of the test tumor cell and the expression profile of a tumor cell of known drug responsivity at least substantially match (*e.g.*, selected sets of genes at elevated levels in the tumor of  
20 known drug responsivity and are also at elevated levels in the test tumor cell), then the drug selected for therapy is the drug to which tumors with that expression pattern respond.

#### Identification of Therapeutic Targets and Anti-Cancer Therapeutic Agents

The present invention also encompasses methods for identification of agents having the ability to modulate activity of a differentially expressed gene product, as well as methods for identifying a  
25 differentially expressed gene product as a therapeutic target for treatment of cancer, especially prostate cancer.

#### Candidate agents

Identification of compounds that modulate activity of a differentially expressed gene product can be accomplished using any of a variety of drug screening techniques. Such agents are candidates  
30 for development of cancer therapies. Of particular interest are screening assays for agents that have tolerable toxicity for normal, non-cancerous human cells. The screening assays of the invention are generally based upon the ability of the agent to modulate an activity of a differentially expressed gene product and/or to inhibit or suppress phenomenon associated with cancer (*e.g.*, cell proliferation, colony formation, cell cycle arrest, metastasis, and the like).

35 The term "agent" as used herein describes any molecule, *e.g.* protein or pharmaceutical, with the capability of modulating a biological activity of a gene product of a differentially expressed gene.

Generally a plurality of assay mixtures are run in parallel with different agent concentrations to obtain a differential response to the various concentrations. Typically, one of these concentrations serves as a negative control, *i.e.* at zero concentration or below the level of detection.

Candidate agents encompass numerous chemical classes, though typically they are organic molecules, preferably small organic compounds having a molecular weight of more than 50 and less than about 2,500 daltons. Candidate agents comprise functional groups necessary for structural interaction with proteins, particularly hydrogen bonding, and typically include at least an amine, carbonyl, hydroxyl or carboxyl group, preferably at least two of the functional chemical groups. The candidate agents often comprise cyclical carbon or heterocyclic structures and/or aromatic or polyaromatic structures substituted with one or more of the above functional groups. Candidate agents are also found among biomolecules including, but not limited to: peptides, saccharides, fatty acids, steroids, purines, pyrimidines, derivatives, structural analogs or combinations thereof.

Candidate agents are obtained from a wide variety of sources including libraries of synthetic or natural compounds. For example, numerous means are available for random and directed synthesis of a wide variety of organic compounds and biomolecules, including expression of randomized oligonucleotides and oligopeptides. Alternatively, libraries of natural compounds in the form of bacterial, fungal, plant and animal extracts (including extracts from human tissue to identify endogenous factors affecting differentially expressed gene products) are available or readily produced. Additionally, natural or synthetically produced libraries and compounds are readily modified through conventional chemical, physical and biochemical means, and may be used to produce combinatorial libraries. Known pharmacological agents may be subjected to directed or random chemical modifications, such as acylation, alkylation, esterification, amidification, *etc.* to produce structural analogs.

Exemplary candidate agents of particular interest include, but are not limited to, antisense polynucleotides, and antibodies, soluble receptors, and the like. Antibodies and soluble receptors are of particular interest as candidate agents where the target differentially expressed gene product is secreted or accessible at the cell-surface (*e.g.*, receptors and other molecule stably-associated with the outer cell membrane).

#### Screening of candidate agents

Screening assays can be based upon any of a variety of techniques readily available and known to one of ordinary skill in the art. In general, the screening assays involve contacting a cancerous cell (preferably a cancerous prostate cell) with a candidate agent, and assessing the effect upon biological activity of a differentially expressed gene product. The effect upon a biological activity can be detected by, for example, detection of expression of a gene product of a differentially expressed gene (*e.g.*, a decrease in mRNA or polypeptide levels, would in turn cause a decrease in biological activity of the gene product). Alternatively or in addition, the effect of the candidate agent

can be assessed by examining the effect of the candidate agent in a functional assay. For example, where the differentially expressed gene product is an enzyme, then the effect upon biological activity can be assessed by detecting a level of enzymatic activity associated with the differentially expressed gene product. The functional assay will be selected according to the differentially expressed gene product. In general, where the differentially expressed gene is increased in expression in a cancerous cell, agents of interest are those that decrease activity of the differentially expressed gene product.

Assays described *infra* can be readily adapted in the screening assay embodiments of the invention. Exemplary assays useful in screening candidate agents include, but are not limited to, hybridization-based assays (*e.g.*, use of nucleic acid probes or primers to assess expression levels), antibody-based assays (*e.g.*, to assess levels of polypeptide gene products), binding assays (*e.g.*, to detect interaction of a candidate agent with a differentially expressed polypeptide, which assays may be competitive assays where a natural or synthetic ligand for the polypeptide is available), and the like. Additional exemplary assays include, but are not necessarily limited to, cell proliferation assays, antisense knockout assays, assays to detect inhibition of cell cycle, assays of induction of cell death/apoptosis, and the like. Generally such assays are conducted *in vitro*, but many assays can be adapted for *in vivo* analyses, *e.g.*, in an animal model of the cancer.

#### Identification of therapeutic targets

In another embodiment, the invention contemplates identification of differentially expressed genes and gene products as therapeutic targets. In some respects, this is the converse of the assays described above for identification of agents having activity in modulating (*e.g.*, decreasing or increasing) activity of a differentially expressed gene product.

In this embodiment, therapeutic targets are identified by examining the effect(s) of an agent that can be demonstrated or has been demonstrated to modulate a cancerous phenotype (*e.g.*, inhibit or suppress or prevent development of a cancerous phenotype). Such agents are generally referred to herein as an "anti-cancer agent", which agents encompass chemotherapeutic agents. For example, the agent can be an antisense oligonucleotide that is specific for a selected gene transcript. For example, the antisense oligonucleotide may have a sequence corresponding to a sequence of a differentially expressed gene described herein, *e.g.*, a sequence of one of SEQ ID NOS:1-2164.

Assays for identification of therapeutic targets can be conducted in a variety of ways using methods that are well known to one of ordinary skill in the art. For example, a test cancerous cell that expresses or overexpresses a differentially expressed gene is contacted with an anti-cancer agent, the effect upon a cancerous phenotype and a biological activity of the candidate gene product assessed. The biological activity of the candidate gene product can be assayed by examining, for example, modulation of expression of a gene encoding the candidate gene product (*e.g.*, as detected by, for example, an increase or decrease in transcript levels or polypeptide levels), or modulation of an enzymatic or other activity of the gene product. The cancerous phenotype can be, for example,

cellular proliferation, loss of contact inhibition of growth (e.g., colony formation), tumor growth (*in vitro* or *in vivo*), and the like. Alternatively or in addition, the effect of modulation of a biological activity of the candidate target gene upon cell death/apoptosis or cell cycle regulation can be assessed.

5 Inhibition or suppression of a cancerous phenotype, or an increase in cell/death apoptosis as a result of modulation of biological activity of a candidate gene product indicates that the candidate gene product is a suitable target for cancer therapy. Assays described infra can be readily adapted in for assays for identification of therapeutic targets. Generally such assays are conducted *in vitro*, but many assays can be adapted for *in vivo* analyses, e.g., in an appropriate, art-accepted animal model of  
10 the cancer.

#### Use of Polynucleotides to Screen for Peptide Analogs and Antagonists

Polypeptides encoded by the instant polynucleotides and corresponding full-length genes can be used to screen peptide libraries to identify binding partners, such as receptors, from among the encoded polypeptides. Peptide libraries can be synthesized according to methods known in the art  
15 (see, e.g., USPN 5,010,175, and WO 91/17823).

Agonists or antagonists of the polypeptides of the invention can be screened using any available method known in the art, such as signal transduction, antibody binding, receptor binding, mitogenic assays, chemotaxis assays, etc. The assay conditions ideally should resemble the conditions under which the native activity is exhibited *in vivo*, that is, under physiologic pH, temperature, and  
20 ionic strength. Suitable agonists or antagonists will exhibit strong inhibition or enhancement of the native activity at concentrations that do not cause toxic side effects in the subject. Agonists or antagonists that compete for binding to the native polypeptide can require concentrations equal to or greater than the native concentration, while inhibitors capable of binding irreversibly to the polypeptide can be added in concentrations on the order of the native concentration.

25 Such screening and experimentation can lead to identification of a novel polypeptide binding partner, such as a receptor, encoded by a gene or a cDNA corresponding to a polynucleotide of the invention, and at least one peptide agonist or antagonist of the novel binding partner. Such agonists and antagonists can be used to modulate, enhance, or inhibit receptor function in cells to which the receptor is native, or in cells that possess the receptor as a result of genetic engineering. Further, if the  
30 novel receptor shares biologically important characteristics with a known receptor, information about agonist/antagonist binding can facilitate development of improved agonists/antagonists of the known receptor.

#### Vaccines and Uses

The differentially expressed nucleic acids and polypeptides produced by the nucleic acids of  
35 the invention can also be used to modulate primary immune response to prevent or treat cancer. Every immune response is a complex and intricately regulated sequence of events involving several cell



types. It is triggered when an antigen enters the body and encounters a specialized class of cells called antigen-presenting cells (APCs). These APCs capture a minute amount of the antigen and display it in a form that can be recognized by antigen-specific helper T lymphocytes. The helper (Th) cells become activated and, in turn, promote the activation of other classes of lymphocytes, such as B cells or cytotoxic T cells. The activated lymphocytes then proliferate and carry out their specific effector functions, which in many cases successfully activate or eliminate the antigen. Thus, activating the immune response to a particular antigen associated with a cancer cell can protect the patient from developing cancer or result in lymphocytes eliminating cancer cells expressing the antigen.

Gene products, including polypeptides, mRNA (particularly mRNAs having distinct secondary and/or tertiary structures), cDNA, or complete gene, can be prepared and used in vaccines for the treatment or prevention of hyperproliferative disorders and cancers. The nucleic acids and polypeptides can be utilized to enhance the immune response, prevent tumor progression, prevent hyperproliferative cell growth, and the like. Methods for selecting nucleic acids and polypeptides that are capable of enhancing the immune response are known in the art. Preferably, the gene products for use in a vaccine are gene products which are present on the surface of a cell and are recognizable by lymphocytes and antibodies.

The gene products may be formulated with pharmaceutically acceptable carriers into pharmaceutical compositions by methods known in the art. The composition is useful as a vaccine to prevent or treat cancer. The composition may further comprise at least one co-immunostimulatory molecule, including but not limited to one or more major histocompatibility complex (MHC) molecules, such as a class I or class II molecule, preferably a class I molecule. The composition may further comprise other stimulator molecules including B7.1, B7.2, ICAM-1, ICAM-2, LFA-1, LFA-3, CD72 and the like, immunostimulatory polynucleotides (which comprise an 5'-CG-3' wherein the cytosine is unmethylated), and cytokines which include but are not limited to IL-1 through IL-15, TNF- $\alpha$ , IFN- $\gamma$ , RANTES, G-CSF, M-CSF, IFN- $\alpha$ , CTAP III, ENA-78, GRO, I-309, PF-4, IP-10, LD-78, MGSA, MIP-1 $\alpha$ , MIP-1 $\beta$ , or combination thereof, and the like for immunopotentiality. In one embodiment, the immunopotentialiators of particular interest are those which facilitate a Th1 immune response.

The gene products may also be prepared with a carrier that will protect the gene products against rapid elimination from the body, such as a controlled release formulation, including implants and microencapsulated delivery systems. Biodegradable polymers can be used, such as ethylene vinyl acetate, polyanhydrides, polyglycolic acid, collagen, polyorthoesters, polylactic acid, and the like. Methods for preparation of such formulations are known in the art.

In the methods of preventing or treating cancer, the gene products may be administered via one of several routes including but not limited to transdermal, transmucosal, intravenous, intramuscular, subcutaneous, intradermal, intraperitoneal, intrathecal, intrapleural, intrauterine, rectal,

vaginal, topical, intratumor, and the like. For transmucosal or transdermal administration, penetrants appropriate to the barrier to be permeated are used in the formulation. Such penetrants are generally known in the art, and include, for example, administration bile salts and fusidic acid derivatives. In addition, detergents may be used to facilitate permeation. Transmucosal administration may be by  
5 nasal sprays or suppositories. For oral administration, the gene products are formulated into conventional oral administration form such as capsules, tablets and toxics.

The gene product is administered to a patient in an amount effective to prevent or treat cancer. In general, it is desirable to provide the patient with a dosage of gene product of at least about 1 pg per Kg body weight, preferably at least about 1 ng per Kg body weight, more preferably at least about  
10 1 µg or greater per Kg body weight of the recipient. A range of from about 1 ng per Kg body weight to about 100 mg per Kg body weight is preferred although a lower or higher dose may be administered. The dose is effective to prime, stimulate and/or cause the clonal expansion of antigen-specific T lymphocytes, preferably cytotoxic T lymphocytes, which in turn are capable of preventing or treating cancer in the recipient. The dose is administered at least once and may be provided as a  
15 bolus or a continuous administration. Multiple administrations of the dose over a period of several weeks to months may be preferable. Subsequent doses may be administered as indicated.

In another method of treatment, autologous cytotoxic lymphocytes or tumor infiltrating lymphocytes may be obtained from a patient with cancer. The lymphocytes are grown in culture, and antigen-specific lymphocytes are expanded by culturing in the presence of the specific gene products  
20 alone or in combination with at least one co-immunostimulatory molecule with cytokines. The antigen-specific lymphocytes are then infused back into the patient in an amount effective to reduce or eliminate the tumors in the patient. Cancer vaccines and their uses are further described in USPN 5,961,978; USPN 5,993,829; USPN 6,132,980; and WO 00/38706.

#### Pharmaceutical Compositions and Uses

25 Pharmaceutical compositions can comprise polypeptides, receptors that specifically bind a polypeptide produced by a differentially expressed gene (e.g., antibodies, or polynucleotides (including antisense nucleotides and ribozymes) of the claimed invention in a therapeutically effective amount. The compositions can be used to treat primary tumors as well as metastases of primary tumors. In addition, the pharmaceutical compositions can be used in conjunction with conventional  
30 methods of cancer treatment, e.g., to sensitize tumors to radiation or conventional chemotherapy.

Where the pharmaceutical composition comprises a receptor (such as an antibody) that specifically binds to a gene product encoded by a differentially expressed gene, the receptor can be coupled to a drug for delivery to a treatment site or coupled to a detectable label to facilitate imaging of a site comprising colon cancer cells. Methods for coupling antibodies to drugs and detectable  
35 labels are well known in the art, as are methods for imaging using detectable labels.

The term "therapeutically effective amount" as used herein refers to an amount of a therapeutic agent to treat, ameliorate, or prevent a desired disease or condition, or to exhibit a detectable therapeutic or preventative effect. The effect can be detected by, for example, chemical markers or antigen levels. Therapeutic effects also include reduction in physical symptoms, such as decreased body temperature.

The precise effective amount for a subject will depend upon the subject's size and health, the nature and extent of the condition, and the therapeutics or combination of therapeutics selected for administration. Thus, it is not useful to specify an exact effective amount in advance. However, the effective amount for a given situation is determined by routine experimentation and is within the judgment of the clinician. For purposes of the present invention, an effective dose will generally be from about 0.01 mg/kg to 50 mg/kg or 0.05 mg/kg to about 10 mg/kg of the DNA constructs in the individual to which it is administered.

A pharmaceutical composition can also contain a pharmaceutically acceptable carrier. The term "pharmaceutically acceptable carrier" refers to a carrier for administration of a therapeutic agent, such as antibodies or a polypeptide, genes, and other therapeutic agents. The term refers to any pharmaceutical carrier that does not itself induce the production of antibodies harmful to the individual receiving the composition, and which can be administered without undue toxicity. Suitable carriers can be large, slowly metabolized macromolecules such as proteins, polysaccharides, polylactic acids, polyglycolic acids, polymeric amino acids, amino acid copolymers, and inactive virus particles. Such carriers are well known to those of ordinary skill in the art. Pharmaceutically acceptable carriers in therapeutic compositions can include liquids such as water, saline, glycerol and ethanol. Auxiliary substances, such as wetting or emulsifying agents, pH buffering substances, and the like, can also be present in such vehicles.

Typically, the therapeutic compositions are prepared as injectables, either as liquid solutions or suspensions; solid forms suitable for solution in, or suspension in, liquid vehicles prior to injection can also be prepared. Liposomes are included within the definition of a pharmaceutically acceptable carrier. Pharmaceutically acceptable salts can also be present in the pharmaceutical composition, e.g., mineral acid salts such as hydrochlorides, hydrobromides, phosphates, sulfates, and the like; and the salts of organic acids such as acetates, propionates, malonates, benzoates, and the like. A thorough discussion of pharmaceutically acceptable excipients is available in Remington's Pharmaceutical Sciences (Mack Pub. Co., N.J. 1991).

#### Delivery Methods

Once formulated, the compositions of the invention can be (1) administered directly to the subject (e.g., as polynucleotide or polypeptides); or (2) delivered ex vivo, to cells derived from the subject (e.g., as in ex vivo gene therapy). Direct delivery of the compositions will generally be accomplished by parenteral injection, e.g., subcutaneously, intraperitoneally, intravenously or

intramuscularly, intratumorally or to the interstitial space of a tissue. Other modes of administration include oral and pulmonary administration, suppositories, and transdermal applications, needles, and gene guns or hyposprays. Dosage treatment can be a single dose schedule or a multiple dose schedule.

Methods for the ex vivo delivery and reimplantation of transformed cells into a subject are known in the art and described in, e.g., WO 93/14778. Examples of cells useful in ex vivo applications include, for example, stem cells, particularly hematopoietic, lymph cells, macrophages, dendritic cells, or tumor cells. Generally, delivery of nucleic acids for both ex vivo and in vitro applications can be accomplished by, for example, dextran-mediated transfection, calcium phosphate precipitation, polybrene mediated transfection, protoplast fusion, electroporation, encapsulation of the polynucleotide(s) in liposomes, and direct microinjection of the DNA into nuclei, all well known in the art.

Once differential expression of a gene corresponding to a polynucleotide of the invention has been found to correlate with a proliferative disorder, such as neoplasia, dysplasia, and hyperplasia, the disorder can be amenable to treatment by administration of a therapeutic agent based on the provided polynucleotide, corresponding polypeptide or other corresponding molecule (e.g., antisense, ribozyme, etc.). In other embodiments, the disorder can be amenable to treatment by administration of a small molecule drug that, for example, serves as an inhibitor (antagonist) of the function of the encoded gene product of a gene having increased expression in cancerous cells relative to normal cells or as an agonist for gene products that are decreased in expression in cancerous cells (e.g., to promote the activity of gene products that act as tumor suppressors).

The dose and the means of administration of the inventive pharmaceutical compositions are determined based on the specific qualities of the therapeutic composition, the condition, age, and weight of the patient, the progression of the disease, and other relevant factors. For example, administration of polynucleotide therapeutic composition agents of the invention includes local or systemic administration, including injection, oral administration, particle gun or catheterized administration, and topical administration. Preferably, the therapeutic polynucleotide composition contains an expression construct comprising a promoter operably linked to a polynucleotide of at least 12, 22, 25, 30, or 35 contiguous nt of the polynucleotide of the invention. Various methods can be used to administer the therapeutic composition directly to a specific site in the body. For example, a small metastatic lesion is located and the therapeutic composition injected several times in several different locations within the body of tumor. Alternatively, arteries that serve a tumor are identified, and the therapeutic composition injected into such an artery, in order to deliver the composition directly into the tumor. A tumor that has a necrotic center is aspirated and the composition injected directly into the now empty center of the tumor. The antisense composition is directly administered to the surface of the tumor, for example, by topical application of the composition. X-ray imaging is used to assist in certain of the above delivery methods.

Targeted delivery of therapeutic compositions containing an antisense polynucleotide, subgenomic polynucleotides, or antibodies to specific tissues can also be used. Receptor-mediated DNA delivery techniques are described in, for example, Findeis et al., *Trends Biotechnol.* (1993) 11:202; Chiu et al., *Gene Therapeutics: Methods And Applications Of Direct Gene Transfer* (J.A. Wolff, ed.) (1994); Wu et al., *J. Biol. Chem.* (1988) 263:621; Wu et al., *J. Biol. Chem.* (1994) 269:542; Zenke et al., *Proc. Natl. Acad. Sci. (USA)* (1990) 87:3655; Wu et al., *J. Biol. Chem.* (1991) 266:338. Therapeutic compositions containing a polynucleotide are administered in a range of about 100 ng to about 200 mg of DNA for local administration in a gene therapy protocol. Concentration ranges of about 500 ng to about 50 mg, about 1 micrograms to about 2 mg, about 5 micrograms to about 500 micrograms, and about 20 micrograms to about 100 micrograms of DNA can also be used during a gene therapy protocol. Factors such as method of action (e.g., for enhancing or inhibiting levels of the encoded gene product) and efficacy of transformation and expression are considerations which will affect the dosage required for ultimate efficacy of the antisense subgenomic polynucleotides.

Where greater expression is desired over a larger area of tissue, larger amounts of antisense subgenomic polynucleotides or the same amounts readministered in a successive protocol of administrations, or several administrations to different adjacent or close tissue portions of, for example, a tumor site, may be required to effect a positive therapeutic outcome. In all cases, routine experimentation in clinical trials will determine specific ranges for optimal therapeutic effect. For polynucleotide related genes encoding polypeptides or proteins with anti-inflammatory activity, suitable use, doses, and administration are described in USP 5,654,173.

The therapeutic polynucleotides and polypeptides of the present invention can be delivered using gene delivery vehicles. The gene delivery vehicle can be of viral or non-viral origin (see generally, Jolly, *Cancer Gene Therapy* (1994) 1:51; Kimura, *Human Gene Therapy* (1994) 5:845; Connelly, *Human Gene Therapy* (1995) 1:185; and Kaplitt, *Nature Genetics* (1994) 6:148). Expression of such coding sequences can be induced using endogenous mammalian or heterologous promoters. Expression of the coding sequence can be either constitutive or regulated.

Viral-based vectors for delivery of a desired polynucleotide and expression in a desired cell are well known in the art. Exemplary viral-based vehicles include, but are not limited to, recombinant retroviruses (see, e.g., WO 90/07936; WO 94/03622; WO 93/25698; WO 93/25234; USP 5,219,740; WO 93/11230; WO 93/10218; USP 4,777,127; GB Patent No. 2,200,651; EP 0 345 242; and WO 91/02805), alphavirus-based vectors (e.g., Sindbis virus vectors, Semliki forest virus (ATCC VR-67; ATCC VR-1247), Ross River virus (ATCC VR-373; ATCC VR-1246) and Venezuelan equine encephalitis virus (ATCC VR-923; ATCC VR-1250; ATCC VR 1249; ATCC VR-532), and adeno-associated virus (AAV) vectors (see, e.g., WO 94/12649, WO 93/03769; WO 93/19191; WO

94/28938; WO 95/11984 and WO 95/00655). Administration of DNA linked to killed adenovirus, as described in Curiel, Hum. Gene Ther. (1992) 3:147, can also be employed.

Non-viral delivery vehicles and methods can also be employed, including, but not limited to, polycationic condensed DNA linked or unlinked to killed adenovirus alone (see, e.g., Curiel, Hum. Gene Ther. (1992) 3:147); ligand-linked DNA (see, e.g., Wu, J. Biol. Chem. (1989) 264:16985); eukaryotic cell delivery vehicles cells (see, e.g., USPN 5,814,482; WO 95/07994; WO 96/17072; WO 95/30763; and WO 97/42338) and nucleic charge neutralization or fusion with cell membranes. Naked DNA can also be employed. Exemplary naked DNA introduction methods are described in WO 90/11092 and USPN 5,580,859. Liposomes that can act as gene delivery vehicles are described in USPN 5,422,120; WO 95/13796; WO 94/23697; WO 91/14445; and EP 0524968. Additional approaches are described in Philip, Mol. Cell Biol. (1994) 14:2411, and in Woffendin, Proc. Natl. Acad. Sci. (1994) 91:1581

Further non-viral delivery suitable for use includes mechanical delivery systems such as the approach described in Woffendin et al., Proc. Natl. Acad. Sci. USA (1994) 91(24):11581. Moreover, the coding sequence and the product of expression of such can be delivered through deposition of photopolymerized hydrogel materials or use of ionizing radiation (see, e.g., USPN 5,206,152 and WO 92/11033). Other conventional methods for gene delivery that can be used for delivery of the coding sequence include, for example, use of hand-held gene transfer particle gun (see, e.g., USPN 5,149,655); use of ionizing radiation for activating transferred gene (see, e.g., USPN 5,206,152 and WO 92/11033).

The present invention will now be illustrated by reference to the following examples which set forth particularly advantageous embodiments. However, it should be noted that these embodiments are illustrative and are not to be construed as restricting the invention in any way.

## EXAMPLES

The following examples are put forth so as to provide those of ordinary skill in the art with a complete disclosure and description of how to make and use the present invention, and are not intended to limit the scope of what the inventors regard as their invention nor are they intended to represent that the experiments below are all or the only experiments performed. It will be readily apparent to those skilled in the art that the formulations, dosages, methods of administration, and other parameters of this invention may be further modified or substituted in various ways without departing from the spirit and scope of the invention. Efforts have been made to ensure accuracy with respect to numbers used (e.g. amounts, temperature, etc.) but some experimental errors and deviations should be accounted for. Unless indicated otherwise, parts are parts by weight, molecular weight is weight average molecular weight, temperature is in degrees Centigrade, and pressure is at or near atmospheric.

Example 1: Source of Biological Materials and Overview of Novel Polynucleotides Expressed by the Biological Materials

Candidate polynucleotides that may represent novel polynucleotides were obtained from cDNA libraries generated from selected cell lines and patient tissues. In order to obtain the candidate polynucleotides, mRNA was isolated from several selected cell lines and patient tissues, and used to construct cDNA libraries. The cells and tissues that served as sources for these cDNA libraries are summarized in Table 1 below.

Human colon cancer cell line Km12L4-A (Morikawa, et al., Cancer Research (1988) 48:6863) is derived from the KM12C cell line. The KM12C cell line (Morikawa et al. Cancer Res. (1988) 48:1943-1948), which is poorly metastatic (low metastatic) was established in culture from a Dukes' stage B2 surgical specimen (Morikawa et al. Cancer Res. (1988) 48:6863). The KM12L4-A is a highly metastatic subline derived from KM12C (Yeatman et al. Nucl. Acids. Res. (1995) 23:4007; Bao-Ling et al. Proc. Annu. Meet. Am. Assoc. Cancer. Res. (1995) 21:3269). The KM12C and KM12C-derived cell lines (e.g., KM12L4, KM12L4-A, etc.) are well-recognized in the art as a model cell line for the study of colon cancer (see, e.g., Moriakawa et al., supra; Radinsky et al. Clin. Cancer Res. (1995) 1:19; Yeatman et al., (1995) supra; Yeatman et al. Clin. Exp. Metastasis (1996) 14:246).

The MDA-MB-231 cell line (Brinkley et al. Cancer Res. (1980) 40:3118-3129) was originally isolated from pleural effusions (Cailleau, J. Natl. Cancer. Inst. (1974) 53:661), is of high metastatic potential, and forms poorly differentiated adenocarcinoma grade II in nude mice consistent with breast carcinoma. The MCF7 cell line was derived from a pleural effusion of a breast adenocarcinoma and is non-metastatic. The MV-522 cell line is derived from a human lung carcinoma and is of high metastatic potential. The UCP-3 cell line is a low metastatic human lung carcinoma cell line; the MV-522 is a high metastatic variant of UCP-3. These cell lines are well-recognized in the art as models for the study of human breast and lung cancer (see, e.g., Chandrasekaran et al., Cancer Res. (1979) 39:870 (MDA-MB-231 and MCF-7); Gastpar et al., J Med Chem (1998) 41:4965 (MDA-MB-231 and MCF-7); Ranson et al., Br J Cancer (1998) 77:1586 (MDA-MB-231 and MCF-7); Kuang et al., Nucleic Acids Res (1998) 26:1116 (MDA-MB-231 and MCF-7); Varki et al., Int J Cancer (1987) 40:46 (UCP-3); Varki et al., Tumour Biol. (1990) 11:327; (MV-522 and UCP-3); Varki et al., Anticancer Res. (1990) 10:637; (MV-522); Kelner et al., Anticancer Res (1995) 15:867 (MV-522); and Zhang et al., Anticancer Drugs (1997) 8:696 (MV522)).

The samples of libraries 15-20 are derived from two different patients (UC#2, and UC#3). The bFGF-treated HMVEC were prepared by incubation with bFGF at 10ng/ml for 2 hrs; the VEGF-treated HMVEC were prepared by incubation with 20ng/ml VEGF for 2 hrs. Following incubation with the respective growth factor, the cells were washed and lysis buffer added for RNA preparation.

GRRpz was derived from normal prostate epithelium. The W0ca cell line is a Gleason Grade 4 cell line.

The source materials for generating the normalized prostate libraries of libraries 25 and 26 were cryopreserved prostate tumor tissue from a patient with Gleason grade 3+3 adenocarcinoma and matched normal prostate biopsies from a pool of at-risk subjects under medical surveillance. The source materials for generating the normalized prostate libraries of libraries 30 and 31 were cryopreserved prostate tumor tissue from a patient with Gleason grade 4+4 adenocarcinoma and matched normal prostate biopsies from a pool of at-risk subjects under medical surveillance.

The source materials for generating the normalized breast libraries of libraries 27, 28 and 29 were cryopreserved breast tissue from a primary breast tumor (infiltrating ductal carcinoma)(library 28), from a lymph node metastasis (library 29), or matched normal breast biopsies from a pool of at-risk subjects under medical surveillance. In each case, prostate or breast epithelia were harvested directly from frozen sections of tissue by laser capture microdissection (LCM, Arcturus Engineering Inc., Mountain View, CA), carried out according to methods well known in the art (see, Simone et al. Am J Pathol. 156(2):445-52 (2000)), to provide substantially homogenous cell samples.

**Table 1. Description of cDNA Libraries**

Library (lib#)	Description	Number of Clones in Library
0	Artificial library composed of deselected clones (clones with no associated variant or cluster)	673
1	Human Colon Cell Line Km12 L4: High Metastatic Potential (derived from Km12C)	308731
2	Human Colon Cell Line Km12C: Low Metastatic Potential	284771
3	Human Breast Cancer Cell Line MDA-MB-231: High Metastatic Potential; micro-mets in lung	326937
4	Human Breast Cancer Cell Line MCF7: Non Metastatic	318979
8	Human Lung Cancer Cell Line MV-522: High Metastatic Potential	223620
9	Human Lung Cancer Cell Line UCP-3: Low Metastatic Potential	312503
12	Human microvascular endothelial cells (HMEC) - UNTREATED (PCR (OligodT) cDNA library)	41938
13	Human microvascular endothelial cells (HMEC) - bFGF TREATED (PCR (OligodT) cDNA library)	42100
14	Human microvascular endothelial cells (HMEC) - VEGF TREATED (PCR (OligodT) cDNA library)	42825
15	Normal Colon - UC#2 Patient (MICRODISSECTED PCR (OligodT) cDNA library)	282722
16	Colon Tumor - UC#2 Patient (MICRODISSECTED PCR (OligodT) cDNA library)	298831
17	Liver Metastasis from Colon Tumor of UC#2 Patient (MICRODISSECTED PCR (OligodT) cDNA library)	303467
18	Normal Colon - UC#3 Patient (MICRODISSECTED PCR (OligodT) cDNA library)	36216
19	Colon Tumor - UC#3 Patient (MICRODISSECTED PCR (OligodT) cDNA library)	41388
20	Liver Metastasis from Colon-Tumor of UC#3 Patient (MICRODISSECTED PCR (OligodT) cDNA library)	30956



Library (lib#)	Description	Number of Clones in Library
21	GRRpz Cells derived from normal prostate epithelium	164801
22	WOca Cells derived from Gleason Grade 4 prostate cancer epithelium	162088
23	Normal Lung Epithelium of Patient #1006 (MICRODISSECTED PCR (OligodT) cDNA library)	306198
24	Primary tumor, Large Cell Carcinoma of Patient #1006 (MICRODISSECTED PCR (OligodT) cDNA library)	309349
25	Normal Prostate Epithelium from Patient IF97-26811	279444
26	Prostate Cancer Epithelium Gleason 3+3 Patient IF97-26811	269406
27	Normal Breast Epithelium from Patient 515	239494
28	Primary Breast tumor from Patient 515	259960
29	Lymph node metastasis from Patient 515	326786
30	Normal Prostate Epithelium from Chiron Patient ID 884	298431
31	Prostate Cancer Epithelium (Gleason 4+4) from Chiron Patient ID 884	331941

#### Characterization of sequences in the libraries

After using the software program Phred (ver 0.000925.c, Green and Weing., ©1993-2000) to select those polynucleotides having the best quality sequence, the polynucleotides were compared against the public databases to identify any homologous sequences. The sequences of the isolated polynucleotides were first masked to eliminate low complexity sequences using the RepeatMasker masking program, publicly available through a web site supported by the University of Washington (See also Smit, A.F.A. and Green, P., unpublished results). Generally, masking does not influence the final search results, except to eliminate sequences of relatively little interest due to their low complexity, and to eliminate multiple "hits" based on similarity to repetitive regions common to multiple sequences, e.g., Alu repeats.

The remaining sequences were then used in a homology search of the GenBank database using the TeraBLAST program (TimeLogic, Crystal Bay, Nevada). TeraBLAST is a version of the publicly available BLAST search algorithm developed by the National Center for Biotechnology, modified to operate at an accelerated speed with increased sensitivity on a specialized computer hardware platform. The program was run with the default parameters recommended by TimeLogic to provide the best sensitivity and speed for searching DNA and protein sequences. Sequences that exhibited greater than 70% overlap, 99% identity, and a p value of less than  $1 \times 10^{-40}$  were discarded. Sequences from this search also were discarded if the inclusive parameters were met, but the sequence was ribosomal or vector-derived.

The resulting sequences from the previous search were classified into three groups (1, 2 and 3 below) and searched in a TeraBLASTX vs. NRP (non-redundant proteins) database search: (1) unknown (no hits in the GenBank search), (2) weak similarity (greater than 45% identity and p value of less than  $1 \times 10^{-5}$ ), and (3) high similarity (greater than 60% overlap, greater than 80% identity,

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and p value less than  $1 \times 10^{-5}$ ). Sequences having greater than 70% overlap, greater than 99% identity, and p value of less than  $1 \times 10^{-40}$  were discarded.

The remaining sequences were classified as unknown (no hits), weak similarity, and high similarity (parameters as above). Two searches were performed on these sequences. First, a TeraBLAST vs. EST database search was performed and sequences with greater than 99% overlap, greater than 99% similarity and a p value of less than  $1 \times 10^{-40}$  were discarded. Sequences with a p value of less than  $1 \times 10^{-65}$  when compared to a database sequence of human origin were also excluded. Second, a TeraBLASTN vs. Patent GeneSeq database was performed and sequences having greater than 99% identity, p value less than  $1 \times 10^{-40}$ , and greater than 99% overlap were discarded.

The remaining sequences were subjected to screening using other rules and redundancies in the dataset. Sequences with a p value of less than  $1 \times 10^{-111}$  in relation to a database sequence of human origin were specifically excluded. The final result provided the sequences listed as SEQ ID NOS:1-1219 in the accompanying Sequence Listing and summarized in Table 2 (inserted prior to claims). Each identified polynucleotide represents sequence from at least a partial mRNA transcript.

#### Summary of polynucleotides of the invention

Table 2 (inserted prior to claims) provides a summary of polynucleotides isolated as described. Specifically, Table 2 provides: 1) the SEQ ID NO ("SEQ ID") assigned to each sequence for use in the present specification; 2) the Cluster Identification No. ("CLUSTER"); 3) the Sequence Name assigned to each sequence; 3) the sequence name ("SEQ NAME") used as an internal identifier of the sequence; 4) the name assigned to the clone from which the sequence was isolated ("CLONE ID"); and 5) the name of the library from which the sequence was isolated ("LIBRARY"). Because at least some of the provided polynucleotides represent partial mRNA transcripts, two or more polynucleotides may represent different regions of the same mRNA transcript and the same gene and/or may be contained within the same clone. Thus, for example, if two or more SEQ ID NOS: are identified as belonging to the same clone, then either sequence can be used to obtain the full-length mRNA or gene. Clones which comprise the sequences described herein were deposited as set out in the tables indicated below (see Example entitled "Deposit Information").

#### Example 2: Contig Assembly

The sequences of the polynucleotides provided in the present invention can be used to extend the sequence information of the gene to which the polynucleotides correspond (e.g., a gene, or mRNA encoded by the gene, having a sequence of the polynucleotide described herein). This expanded sequence information can in turn be used to further characterize the corresponding gene, which in turn provides additional information about the nature of the gene product (e.g., the normal function of the gene product). The additional information can serve to provide additional evidence of the gene

product's use as a therapeutic target, and provide further guidance as to the types of agents that can modulate its activity.

For example, a contig was assembled using the sequence of a polynucleotide described herein.

A "contig" is a contiguous sequence of nucleotides that is assembled from nucleic acid sequences

5 having overlapping (e.g., shared or substantially similar) sequence information. The sequences of publicly-available ESTs (Expressed Sequence Tags) and the sequences of various of the above-described polynucleotides were used in the contig assembly. The contig was assembled using the software program Sequencher, version 4.05, according to the manufacturer's instructions. The sequence information obtained in the contig assembly was then used to obtain a consensus sequence  
10 derived from the contig using the Sequencher program. The resulting consensus sequence was used to search both the public databases as well as databases internal to the applicants to match the consensus polynucleotide with homology data and/or differential gene expressed data.

The final result provided the sequences listed as SEQ ID NOS: 1220-1428 in the accompanying Sequence Listing and summarized in Tables 3 and 4 (inserted prior to claims). Table 3  
15 provides a summary of the consensus sequences assembled as described. Specifically, Table 3 provides: 1) the SEQ ID NO ("SEQ ID") assigned to each consensus sequence for use in the present specification; 2) the Cluster Identification No. ("CLUSTER"); and 3) the consensus sequence name ("CONSENSUS SEQ NAME") used as an internal identifier of the sequence.

A correlation between the polynucleotide used in consensus sequence assembly as described  
20 above and the corresponding consensus sequence is contained in Table 4. Specifically Table 4 provides: 1) the SEQ ID NO of the consensus sequence ("CONSENSUS SEQ ID"); 2) the consensus sequence name ("CONSENSUS SEQ NAME") used as an internal identifier of the sequence; 3) the SEQ ID NO of the polynucleotide ("POLYNTD SEQ ID") of SEQ ID NOS: 1-1219 used in assembly of the consensus sequence; and 4) the sequence name ("POLYNTD SEQ NAME") of the  
25 polynucleotide of SEQ ID NOS: 1-1219 used in assembly of the consensus sequence.

#### Example 3: Additional Gene Characterization

Sequences of the polynucleotides of SEQ ID NOS: 1-1219 were used as a query sequence in a TeraBLASTN search of the DoubleTwist Human Genome Sequence Database (DoubleTwist, Inc., Oakland, CA), which contains all the human genomic sequences that have been assembled into a  
30 contiguous model of the human genome. Predicted cDNA and protein sequences were obtained where a polynucleotide of the invention was homologous to a predicted full-length gene sequence. Alternatively, a sequence of a contig or consensus sequence described herein could be used directly as a query sequence in a TeraBLASTN search of the DoubleTwist Human Genome Sequence Database.

The final results of the search provided the predicted cDNA sequences listed as SEQ ID NOS:  
35 1429-1485 in the accompanying Sequence Listing and summarized in Table 5 (inserted prior to claims), and the predicted protein sequences listed as SEQ ID NOS: 1486-1542 in the accompanying

Sequence Listing and summarized in Table 6 (inserted prior to claims). Specifically, Table 5 provides: 1) the SEQ ID NO ("SEQ ID") assigned to each cDNA sequence for use in the present specification; 2) the cDNA sequence name ("cDNA SEQ NAME") used as an internal identifier of the sequence; 3) the chromosome ("CHROM") containing the gene corresponding to the cDNA sequence; and 4) the exon ("EXON") of the gene corresponding to the cDNA sequence to which the polynucleotide of SEQ ID NOS: 1-1219 maps. Table 6 provides: 1) the SEQ ID NO ("SEQ ID") assigned to each protein sequence for use in the present specification; 2) the protein sequence name ("PROTEIN SEQ NAME") used as an internal identifier of the sequence; 3) the chromosome ("CHROM") containing the gene corresponding to the cDNA sequence; and 4) the exon ("EXON") of the gene corresponding to the cDNA and protein sequence to which the polynucleotide of SEQ ID NOS: 1-1219 maps.

A correlation between the polynucleotide used as a query sequence as described above and the corresponding predicted cDNA and protein sequences is contained in Table 7. Specifically Table 7 provides: 1) the SEQ ID NO of the cDNA ("cDNA SEQ ID"); 2) the cDNA sequence name ("cDNA SEQ NAME") used as an internal identifier of the sequence; 3) the SEQ ID NO of the protein ("PROTEIN SEQ ID") encoded by the cDNA sequence 4) the sequence name of the protein ("PROTEIN SEQ NAME") encoded by the cDNA sequence; 5) the SEQ ID NO of the polynucleotide ("POLYNTD SEQ ID") of SEQ ID NOS: 1-1219 that maps to the cDNA and protein; and 6) the sequence name ("POLYNTD SEQ NAME") of the polynucleotide of SEQ ID NOS: 1-1219 that maps to the cDNA and protein.

Through contig and consensus sequence assembly and the use of homology searching software programs, the sequence information provided herein can be readily extended to confirm, or confirm a predicted, gene having the sequence of the polynucleotides described in the present invention. Further the information obtained can be used to identify the function of the gene product of the gene corresponding to the polynucleotides described herein. While not necessary to the practice of the invention, identification of the function of the corresponding gene, can provide guidance in the design of therapeutics that target the gene to modulate its activity and modulate the cancerous phenotype (e.g., inhibit metastasis, proliferation, and the like).

#### Example 4: Results of Public Database Search to Identify Function of Gene Products

SEQ ID NOS: 1-1485 were translated in all three reading frames, and the nucleotide sequences and translated amino acid sequences used as query sequences to search for homologous sequences in the GenBank (nucleotide sequences) database. Query and individual sequences were aligned using the TeraBLAST program available from TimeLogic, Crystal Bay, Nevada. The sequences were masked to various extents to prevent searching of repetitive sequences or poly-A sequences, using the RepeatMasker masking program for masking low complexity as described above.

Table 8 (inserted prior to claims) provides the alignment summaries having a p value of  $1 \times 10^{-2}$  or less indicating substantial homology between the sequences of the present invention and those of the indicated public databases. Specifically, Table 8 provides: 1) the SEQ ID NO ("SEQ ID") of the query sequence; 2) the sequence name ("SEQ NAME") used as an internal identifier of the query sequence; 3) the accession number ("ACCESSION") of the GenBank database entry of the homologous sequence; 4) a description of the GenBank sequences ("GENBANK DESCRIPTION"); and 5) the score of the similarity of the polynucleotide sequence and the GenBank sequence ("GENBANK SCORE"). The alignments provided in Table 8 are the best available alignment to a DNA sequence at a time just prior to filing of the present specification. Incorporated by reference is all publicly available information regarding the sequence listed in Table 8 and their related sequences. The search program and database used for the alignment, as well as the calculation of the p value are also indicated. Full length sequences or fragments of the polynucleotide sequences can be used as probes and primers to identify and isolate the full length sequence of the corresponding polynucleotide.

#### Example 5: Members of Protein Families

SEQ ID NOS:1-1219 were used to conduct a profile search as described in the specification above. Several of the polynucleotides of the invention were found to encode polypeptides having characteristics of a polypeptide belonging to a known protein family (and thus represent members of these protein families) and/or comprising a known functional domain. Table 9 (inserted prior to claims) provides: 1) the SEQ ID NO ("SEQ ID") of the query polynucleotide sequence; 2) the sequence name ("SEQ NAME") used as an internal identifier of the query sequence; 3) the name ("PFAM NAME") of the profile hit; 4) a brief description of the profile hit ("PFAM DESCRIPTION"); 5) the score ("SCORE") of the profile hit; 6) the starting nucleotide of the profile hit ("START"); and 7) the ending nucleotide of the profile hit ("END").

In addition, SEQ ID NOS:1486-1542 were also used to conduct a profile search as described above. Several of the polypeptides of the invention were found to have characteristics of a polypeptide belonging to a known protein family (and thus represent members of these protein families) and/or comprising a known functional domain. Table 10 (inserted prior to claims) provides: 1) the SEQ ID NO ("SEQ ID") of the query protein sequence; 2) the sequence name ("PROTEIN SEQ NAME") used as an internal identifier of the query sequence; 3) the name ("PFAM NAME") of the profile hit; 4) a brief description of the profile hit ("PFAM DESCRIPTION"); 5) the score ("SCORE") of the profile hit; 6) the starting residue of the profile hit ("START"); and 7) the ending residue of the profile hit ("END").

Some SEQ ID NOS exhibited multiple profile hits where the query sequence contains overlapping profile regions, and/or where the sequence contains two different functional domains. Each of the profile hits of Tables 9 and 10 is described in more detail below. The acronyms for the

profiles (provided in parentheses) are those used to identify the profile in the Pfam, Prosite, and InterPro databases. The Pfam database can be accessed through web sites supported by Genome Sequencing Center at the Washington University School of Medicine or by the European Molecular Biology Laboratories in Heidelberg, Germany. The Prosite database can be accessed at the Expasy Molecular Biology Server on the internet. The InterPro database can be accessed at a web site supported by the EMBL European Bioinformatics Institute. The public information available on the Pfam, Prosite, and InterPro databases regarding the various profiles, including but not limited to the activities, function, and consensus sequences of various proteins families and protein domains, is incorporated herein by reference.

Epidermal Growth Factor (EGF; Pfam Accession No. PF00008). SEQ ID NOS:417 and 418 represent polynucleotides encoding a member of the EGF family of proteins. The distinguishing characteristic of this family is the presence of a sequence of about thirty to forty amino acid residues found in epidermal growth factor (EGF) which has been shown to be present, in a more or less conserved form, in a large number of other proteins (Davis, *New Biol.* (1990) 2:410-419; Blomquist *et al.*, *Proc. Natl. Acad. Sci. U.S.A.* (1984) 81:7363-7367; Barkert *et al.*, *Protein Nucl. Acid Enz.* (1986) 29:54-86; Doolittle *et al.*, *Nature.* (1984) 307:558-560; Appella *et al.*, *FEBS Lett.* (1988) 231:1-4; Campbell and Bork, *Curr. Opin. Struct. Biol.* (1993) 3:385-392). A common feature of the domain is that the conserved pattern is generally found in the extracellular domain of membrane-bound proteins or in proteins known to be secreted. The EGF domain includes six cysteine residues which have been shown to be involved in disulfide bonds. The main structure is a two-stranded beta-sheet followed by a loop to a C-terminal short two-stranded sheet. Subdomains between the conserved cysteines strongly vary in length. These consensus patterns are used to identify members of this family: C-x-C-x(5)-G-x(2)-C and C-x-C-x(s)-[GP]-[FYW]-x(4,8)-C.

Seven Transmembrane Integral Membrane Proteins -- Rhodopsin Family (7tm 1; Pfam Accession No. PF00001). SEQ ID NO:321 corresponds to a sequence encoding a polypeptide that is a member of the seven transmembrane (7tm) receptor rhodopsin family. G-protein coupled receptors of the (7tm) rhodopsin family (also called R7G) are an extensive group of hormones, neurotransmitters, and light receptors which transduce extracellular signals by interaction with guanine nucleotide-binding (G) proteins (Strosberg, *Eur. J. Biochem.* (1991) 196:1; Kerlavage, *Curr. Opin. Struct. Biol.* (1991) 1:394; Probst *et al.*, *DNA Cell Biol.* (1992) 11:1; Savarese *et al.*, *Biochem. J.* (1992) 283:1. The consensus pattern that contains the conserved triplet and that also spans the major part of the third transmembrane helix is used to detect this widespread family of proteins: [GSTALIVMFYWC]-[GSTANCPDE]-{EDPKRH}-x(2)-[LIVMNQGA]-x(2)-[LIVMFT]-[GSTANC]-[LIVMFYWTAC]-[DENH]-R-[FYWCSH]-x(2)-[LIVM].

Basic Region Plus Leucine Zipper Transcription Factors (bZIP; Pfam Accession No. PF00170). SEQ ID NO:638 represents a polynucleotide encoding a novel member of the family

of basic region plus leucine zipper transcription factors. The bZIP superfamily (Hurst, *Protein Prof.* (1995) 2:105; and Ellenberger, *Curr. Opin. Struct. Biol.* (1994) 4:12) of eukaryotic DNA-binding transcription factors encompasses proteins that contain a basic region mediating sequence-specific DNA-binding followed by a leucine zipper required for dimerization. The consensus pattern for this protein family is: [KR]-x(1,3)-[RKSAQ]-N-x(2)-[SAQ](2)-x-[RKTAENQ]-x-R-x-[RK].

Reverse Transcriptase (rvt; Pfam Accession No. PF00078). SEQ ID NO:137 represents a polynucleotide encoding a reverse transcriptase, which occurs in a variety of mobile elements, including retrotransposons, retroviruses, group II introns, bacterial msDNAs, hepadnaviruses, and caulimoviruses (Xiong and Eickbush, *EMBO J* (1990) 9:3353-3362). Reverse transcriptases catalyze RNA-template-directed extension of the 3'-end of a DNA strand by one deoxynucleotide at a time and require an RNA or DNA primer.

KRAB box (KRAB; Pfam Accession No. PF01352). SEQ ID NO:1012 represents a polypeptide having a Krueppel-associated box (KRAB). A KRAB box is a domain of around 75 amino acids that is found in the N-terminal part of about one third of eukaryotic Krueppel-type C2H2 zinc finger proteins (ZFPs). It is enriched in charged amino acids and can be divided into subregions A and B, which are predicted to fold into two amphipathic alpha-helices. The KRAB A and B boxes can be separated by variable spacer segments and many KRAB proteins contain only the A box.

The KRAB domain functions as a transcriptional repressor when tethered to the template DNA by a DNA-binding domain. A sequence of 45 amino acids in the KRAB A subdomain has been shown to be necessary and sufficient for transcriptional repression. The B box does not repress by itself but does potentiate the repression exerted by the KRAB A subdomain. Gene silencing requires the binding of the KRAB domain to the RING-B box-coiled coil (RBCC) domain of the KAP-1/ITF1-beta corepressor. As KAP-1 binds to the heterochromatin proteins HP1, it has been proposed that the KRAB-ZFP-bound target gene could be silenced following recruitment to heterochromatin.

KRAB-ZFPs constitute one of the single largest class of transcription factors within the human genome, and appear to play important roles during cell differentiation and development. The KRAB domain is generally encoded by two exons. The regions coded by the two exons are known as KRAB-A and KRAB-B.

Armadillo/beta-catenin-like repeat (Armadillo\_seg; Pfam Accession No. PF00514). SEQ ID NO: 1486 represents a polypeptide having sequence similarity with the armadillo/beta-catenin-like repeat (armadillo). The armadillo repeat is an approximately 40 amino acid long tandemly repeated sequence motif first identified in the *Drosophila* segment polarity gene armadillo. Similar repeats were later found in the mammalian armadillo homolog beta-catenin, the junctional plaque protein plakoglobin, the adenomatous polyposis coli (APC) tumor suppressor protein, and a number of other proteins (Peifer *et al.*, *Cell* 76(2):786-791 (1994)).

The 3 dimensional fold of an armadillo repeat is known from the crystal structure of beta-catenin (Rojas *et al.*, *Cell* 95:105-130 (1998)). There, the 12 repeats form a superhelix of alpha-helices, with three helices per unit. The cylindrical structure features a positively charged groove which presumably interacts with the acidic surfaces of the known interaction partners of beta-catenin.

- 5        Cadherin domain (cadherin: Pfam Accession No. PF00028). SEQ ID NO: 1523 represents a polypeptide having sequence similarity to a cadherin domain. Cadherins are a family of animal glycoproteins responsible for calcium-dependent cell-cell adhesion (Takeichi, *Annu. Rev. Biochem.* 59:237-252(1990); Takeichi, *Trends Genet.* 3:213-217(1987)). Cadherins preferentially interact with themselves in a homophilic manner in connecting cells; thus acting as both receptor and ligand. A
- 10    wide number of tissue-specific forms of cadherins are known, for example: Epithelial (E-cadherin) (CDH1); Neural (N-cadherin) (CDH2); Placental (P-cadherin) (CDH3); Retinal (R-cadherin) (CDH4); Vascular endothelial (VE-cadherin) (CDH5); Kidney (K-cadherin) (CDH6); Cadherin-8 (CDH8); Cadherin-9 (CDH9); Osteoblast (OB-cadherin) (CDH11); Brain (BR-cadherin) (CDH12); T-cadherin (truncated cadherin) (CDH13); Muscle (M-cadherin) (CDH15); Kidney (Ksp-cadherin) (CDH16); and
- 15    Liver-intestine (LI-cadherin) (CDH17).

- 20    Structurally, cadherins are built of the following domains: a signal sequence, followed by a propeptide of about 130 residues, then an extracellular domain of around 600 residues, then a transmembrane region, and finally a C-terminal cytoplasmic domain of about 150 residues. The extracellular domain can be sub-divided into five parts: there are four repeats of about 110 residues followed by a region that contains four conserved cysteines. The calcium-binding region of cadherins may be located in the extracellular repeats. The signature pattern for the repeated domain is located in the C-terminal extremity, which is its best conserved region. The pattern includes two conserved aspartic acid residues and two asparagines; these residues could be implicated in the binding of calcium. The consensus pattern is: [LIV]-x-[LIV]-x-D-x-N-D-[NH]-x-P.

- 25    CBS domain (CBS: Pfam Accession No. PF00571). SEQ ID NOS:1510 and 1511 represent polypeptides having sequence similarity to CBS domains, which are present in all 3 forms of cellular life, including two copies in inosine monophosphate dehydrogenase, of which one is disordered in the crystal structure. A number of disease states are associated with CBS-containing proteins including homocystinuria, Becker's and Thomsen disease.

- 30    CBS domains are small intracellular modules of unknown function. They are mostly found in 2 or four copies within a protein. Pairs of CBS domains dimerise to form a stable globular domain (Zhang *et al.*, *Biochemistry* 38:4691-4700 (1999)). Two CBS domains are found in inosine-monophosphate dehydrogenase from all species, however the CBS domains are not needed for activity. CBS domains are found attached to a wide range of other protein domains suggesting that
- 35    CBS domains may play a regulatory role. The region containing the CBS domains in Cystathionine-beta synthase is involved in regulation by S-AdoMet (Zhang *et al.*, *Biochemistry* 38:4691-4700



(1999)). The 3D Structure is found as a sub-domain in TIM barrel of inosine-monophosphate dehydrogenase.

Phorbol esters/diacylglycerol binding domain (C1 domain) (DAG\_PE-bind; Pfam Accession No. PF00130). SEQ ID NO: 1514 represents a polypeptide having sequence similarity to the Phorbol esters/diacylglycerol binding domain (C1 domain). Diacylglycerol (DAG) is an important second messenger. Phorbol esters (PE) are analogues of DAG and potent tumor promoters that cause a variety of physiological changes when administered to both cells and tissues. DAG activates a family of serine/threonine protein kinases, collectively known as protein kinase C (PKC) (Azzi *et al.*, *Eur. J. Biochem.* 208:547-557 (1992)). Phorbol esters can also directly stimulate PKC.

The N-terminal region of PKC, known as C1, has been shown to bind PE and DAG in a phospholipid and zinc-dependent fashion(Ono *et al.*, *Proc. Natl. Acad. Sci. U.S.A.* 86:4868-4871 (1989)). The C1 region contains one or two copies (depending on the isozyme of PKC) of a cysteine-rich domain about 50 amino-acid residues long and essential for DAG/PE-binding. The DAG/PE-binding domain binds two zinc ions; the ligands of these metal ions are probably the six cysteines and two histidines that are conserved in the C1 domain. The consensus sequence for the C1 domain is: H-x-[LIVMFYW]-x(8,11)-C-x(2)-C-x(3)-[LIVMFC]-x(5,10)-C-x(2)-C-x(4)-[HD]-x(2)-C-x(5,9)-C [All the C and H are involved in binding Zinc].

GATA zinc finger (GATA; Pfam Accession No. PF00320). SEQ ID NO:1520 represents a polypeptide having sequence similarity to GATA zinc finger. A number of transcription factors, including erythroid-specific transcription factor and nitrogen regulatory proteins, specifically bind the DNA sequence (A/T)GATA(A/G) in the regulatory regions of genes (Yamamoto *et al.*, *Genes Dev.* 4:1650-1662 (1990)) and are consequently termed GATA-binding transcription factors. The interactions occur via highly-conserved zinc finger domains in which the zinc ion is coordinated by 4 cysteine residues (Evans and Felsenfeld, *Cell* 58:877-885 (1989); Omichinski *et al.*, *Science* 261:438-446 (1993)).

NMR studies have shown the core of the zinc finger to comprise 2 irregular anti-parallel beta-sheets and an alpha-helix, followed by a long loop to the C-terminal end of the finger. The N-terminal part, which includes the helix, is similar in structure, but not sequence, to the N-terminal zinc module of the glucocorticoid receptor DNA-binding domain. The helix and the loop connecting the 2 beta-sheets interact with the major groove of the DNA, while the C-terminal tail wraps around into the minor groove. It is this tail that is the essential determinant of specific binding. Interactions between the zinc finger and DNA are mainly hydrophobic, explaining the preponderance of thymines in the binding site; a large number of interactions with the phosphate backbone have also been observed (Omichinski *et al.*, *Science* 261:438-446 (1993)). Two GATA zinc fingers are found in the GATA transcription factors; however, there are several proteins which only contains a single copy of the

domain. The consensus sequence of the domain is: C-x-[DN]-C-x(4,5)-[ST]-x(2)-W-[HR]-[RK]-x(3)-[GN]-x(3,4)-C-N-[AS]-C [The four C's are zinc ligands].

5 Glutathione S-transferase, N-terminal domain (GST\_N; Pfam Accession No. PF02798), SEQ ID NO: 1507 represents a polypeptide having sequence similarity to Glutathione S-transferase, N-terminal domain. In eukaryotes, glutathione S-transferases (GSTs) participate in the detoxification of reactive electrophilic compounds by catalysing their conjugation to glutathione. The GST domain is also found in S-crystallins from squid, and proteins with no known GST activity, such as eukaryotic elongation factors 1-gamma and the HSP26 family of stress-related proteins, which include auxin-regulated proteins in plants and stringent starvation proteins in E. coli. The major lens polypeptide of Cephalopoda is also a GST.

Bacterial GSTs of known function often have a specific, growth-supporting role in biodegradative metabolism: epoxide ring opening and tetrachlorohydroquinone reductive dehalogenation are two examples of the reactions catalysed by these bacterial GSTs. Some regulatory proteins, like the stringent starvation proteins, also belong to the GST family. GST seems to be absent from Archaea in which gamma-glutamylcysteine substitute to glutathione as major thiol.

Glutathione S-transferases form homodimers, but in eukaryotes can also form heterodimers of the A1 and A2 or YC1 and YC2 subunits. The homodimeric enzymes display a conserved structural fold. Each monomer is composed of a distinct N-terminal sub-domain, which adopts the thioredoxin fold, and a C-terminal all-helical sub-domain.

20 GTF2I-like repeat (GTF2I; Pfam Accession No. PF02946), SEQ ID NOS:1500, 1501, and 1542 represent polypeptides having sequence similarity to proteins having GTF2I-like repeat. This region of sequence similarity is found up to six times in a variety of proteins including GTF2I. It has been suggested that this may be a DNA binding domain (O'Mahoney *et al.*, *Mol. Cell. Biol.* 18:6641-6652 (1998); Osborne *et al.*, *Genomics* 57:279-284 (1999)).

25 Core histone H2A/H2B/H3/H4 (histone; Pfam Accession No. PF00125), SEQ ID NO:1497 represents a polypeptide having sequence similarity to core histone H2A/H2B/H3/H4 family polypeptides. Histone H2A is one of the four histones, along with H2B, H3 and H4, which forms the eukaryotic nucleosome core. Using alignments of histone H2A sequences (Wells and Brown, *Nucleic Acids Res.* 19:2173-2188(1991); Thatcher and Gorovsky, *Nucleic Acids Res.* 22:174-179(1994)) a conserved region in the N-terminal part of H2A was used to develop a signature pattern. This region is conserved both in classical S-phase regulated H2A's and in variant histone H2A's which are synthesized throughout the cell cycle. The consensus pattern is: [AC]-G-L-x-F-P-V.

Histone H4, along with H3, plays a central role in nucleosome formation. The sequence of histone H4 has remained almost invariant in more than 2 billion years of evolution (Thatcher and Gorovsky, *Nucleic Acids Res.* 22:174-179(1994)). The region used as a signature pattern is a pentapeptide found in positions 14 to 18 of all H4 sequences. It contains a lysine residue which is

often acetylated (Doenecke and Gallwitz, *Mol. Cell. Biochem.* 44:113-128(1982)) and a histidine residue which is implicated in DNA-binding (Ebralidse *et al.*, *Nature* 331:365-367(1988)). The consensus pattern is: G-A-K-R-H.

Histone H3 is a highly conserved protein of 135 amino acid residues (Wells and Brown, *Nucleic Acids Res.* 19:2173-2188(1991); Thatcher and Gorovsky, *Nucleic Acids Res.* 22:174-179(1994)). Two signature patterns have been developed, the first one corresponds to a perfectly conserved heptapeptide in the N-terminal part of H3, while the second one is derived from a conserved region in the central section of H3. The consensus patterns are: K-A-P-R-K-Q-L and P-F-x-[RA]-L-[VA]-[KRQ]-[DEG]-[IV].

The signature pattern of histone H2B corresponds to a conserved region in the C-terminal part of the protein. The consensus pattern is: [KR]-E-[LIVM]-[EQ]-T-x(2)-[KR]-x-[LIVM](2)-x-[PAG]-[DE]-L-x-[KR]-H-A-[LIVM]-[STA]-E-G

HMG (high mobility group) box (HMG\_box; Pfam Accession No. PF00505). SEQ ID NO:1525 corresponds to a polypeptide having sequence similarity to high mobility group proteins, a family of relatively low molecular weight non-histone components in chromatin. HMG1 (also called HMG-T in fish) and HMG2 (Bustin *et al.*, *Biochim. Biophys. Acta* 1049: 231-243(1990)) are two highly related proteins that bind single-stranded DNA preferentially and unwind double-stranded DNA. HMG1/2 have about 200 amino acid residues with a highly acidic C-terminal section which is composed of an uninterrupted stretch of from 20 to 30 aspartic and glutamic acid residues; the rest of the protein sequence is very basic. In addition to the HMG1 and HMG2 proteins, HMG-domains occur in single or multiple copies in the following protein classes; the SOX family of transcription factors; SRY sex determining region Y protein and related proteins; LEF1 lymphoid enhancer binding factor 1; SSRP recombination signal recognition protein; MTF1 mitochondrial transcription factor 1; UBF1/2 nucleolar transcription factors; Abf2 yeast ARS-binding factor; and yeast transcription factors Ixr1, Rox1, Nhp6a, Nhp6b and Spp41.

Importin beta binding domain (IBB; Pfam Accession No. PF01749). SEQ ID NO: 1486 represents a polypeptide having sequence similarity to importin beta binding domain family polypeptides. This family consists of the importin alpha (karyopherin alpha), importin beta (karyopherin beta) binding domain. The domain mediates formation of the importin alpha beta complex; required for classical NLS import of proteins into the nucleus, through the nuclear pore complex and across the nuclear envelope. Also in the alignment is the NLS of importin alpha which overlaps with the IBB domain (Moroianu *et al.*, *Proc. Natl. Acad. Sci. U.S.A.* 93:6572-6576(1996)).

T-box domain (T-box; Pfam Accession No. PF00907). SEQ ID NOS:1518 represents a polypeptide having sequence similarity to proteins having a T-box domain. The T-box gene family is an ancient group of putative transcription factors that appear to play a critical role in the development of all animal species. These genes were uncovered on the basis of similarity to the DNA binding

domain (Papaioannou and Silver, *Bioessays* 20:9-19 (1998)) of murine Brachyury (T) gene product, which similarity is the defining feature of the family. The Brachyury gene is named for its phenotype, which was identified 70 years ago as a mutant mouse strain with a short blunted tail. The gene, and its paralogs, have become a well-studied model for the family, and hence much of what is known about the T-box family is derived from the murine Brachyury gene.

Consistent with its nuclear location, Brachyury protein has a sequence-specific DNA-binding activity and can act as a transcriptional regulator (Wattler *et al.*, *Genomics* 48:24-33(1998)). Homozygous mutants for the gene undergo extensive developmental anomalies, thus rendering the mutation lethal (Kavka and Green, *Biochim. Biophys. Acta* 1333(2) (1997)). The postulated role of Brachyury is as a transcription factor, regulating the specification and differentiation of posterior mesoderm during gastrulation in a dose-dependent manner (Papaioannou and Silver, *Bioessays* 20:9-19 (1998)).

Common features shared by T-box family members are, DNA-binding and transcriptional regulatory activity, a role in development and conserved expression patterns. Most of the known genes in all species are expressed in mesoderm or mesoderm precursors (Papaioannou, *Trends Genet.* 13:212-213(1997)). Members of the T-box family contain a domain of about 170 to 190 amino acids known as the T-box domain (Papaioannou, *Trends Genet.* 13: 212-213(1997); Bollag *et al.*, *Nat. Genet.* 7: 383-389(1994); Agulnik *et al.*, *Genetics* 144:249-254(1996)) and which probably binds DNA. As signature patterns for the T-domain, we selected two conserved regions. The first region corresponds to the N-terminal of the domain and the second one to the central part. The consensus sequences are: L-W-x(2)-[FC]-x(3,4)-[NT]-E-M-[LIV](2)-T-x(2)-G-[RG]-[KRQ] and [LIVMFYW]-H-[PADH]-[DENQ]-[GS]-x(3)-G-x(2)-W-M-x(3)-[IVA]-x-F.

60s Acidic ribosomal protein (60s ribosomal; Pfam Accession No. PF00428). SEQ ID NO: 905 represents a polynucleotide encoding a member of the 60s acidic ribosomal protein family. The 60S acidic ribosomal protein plays an important role in the elongation step of protein synthesis. This family includes archaeobacterial L12, eukaryotic P0, P1 and P2 (Remacha *et al.*, *Biochem. Cell Biol.* 73:959-968(1995)).

Some of the proteins in this family are allergens. A nomenclature system has been established for antigens (allergens) that cause IgE-mediated atopic allergies in humans (WHO/IUIS Allergen Nomenclature Subcommittee King T.P., Hoffmann D., Loewenstein H., Marsh D.G., Platts-Mills T.A.E., Thomas W. Bull. World Health Organ. 72:797-806(1994)). This nomenclature system is defined by a designation that is composed of the first three letters of the genus; a space; the first letter of the species name; a space and an arabic number. In the event that two species names have identical designations, they are discriminated from one another by adding one or more letters (as necessary) to each species designation. The allergens in this family include allergens with the following designations: Alt a 6, Alt a 12, Cla h 3, Cla h 4, and Cla h 12.

AP endonuclease family 1 (AP\_endonuclease1; Pfam Accession No. PF01260). SEQ ID NOS:358 and 836 correspond to a polynucleotide encoding a member of the family of polypeptides designated AP endonuclease family 1. DNA damaging agents such as the antitumor drugs bleomycin and neocarzinostatin or those that generate oxygen radicals produce a variety of lesions in DNA.

- 5 Amongst these is base-loss which forms apurinic/aprimidinic (AP) sites or strand breaks with atypical 3'-termini. DNA repair at the AP sites is initiated by specific endonuclease cleavage of the phosphodiester backbone. Such endonucleases are also generally capable of removing blocking groups from the 3'-terminus of DNA strand breaks.

- AP endonucleases can be classified into two families on the basis of sequence similarity. This family contains members of AP endonuclease family 1. Except for Rrp1 and arp, these enzymes are proteins of about 300 amino-acid residues. Rrp1 and arp both contain additional and unrelated sequences in their N-terminal section (about 400 residues for Rrp1 and 270 for arp). The proteins contain glutamate which has been shown (Mol *et al.*, *Nature* 374: 381-386(1995)), in the *Escherichia coli* enzyme to bind a divalent metal ion such as magnesium or manganese. The consensus sequences for this family of polypeptides are: [APF]-D-[LIVMF](2)-x-[LIVM]-Q-E-x-K [E binds a divalent metal ion]; D-[ST]-[FY]-R-[KH]-x(7,8)-[FYW]-[ST]-[FYW](2); and N-x-G-x-R-[LIVM]-D-[LIVMFYH]-x-[LV]-x-S

- Bowman-Birk serine protease inhibitor family (Bowman-Birk\_leg; Pfam Accession No. 00228). SEQ ID NO: 321 represents a polynucleotide encoding a polypeptide having sequence similarity to a member of the Bowman-Birk serine protease inhibitor family. The Bowman-Birk inhibitor family (Laskowski and Kato, *Annu. Rev. Biochem.* 49:593-626(1980)) is one of the numerous families of serine proteinase inhibitors and has a duplicated structure and generally possesses two distinct inhibitory sites.

- These inhibitors are found in the seeds of all leguminous plants as well as in cereal grains. In cereals they exist in two forms, one of which is a duplication of the basic structure (Tashiro *et al.*, *J. Biochem.* 102:297-306(1987)). The signature pattern for sequences belonging to this family of inhibitors is in the central part of the domain and includes four cysteines. The consensus pattern is: C-x(5,6)-[DENQKRHSTA]-C-[PASTDH]-[PASTDK]-[ASTDV]-C-[NDEKS]-[DEKRHSTA]-C [The four C's are involved in disulfide bonds]. Note that this pattern can be found twice in some duplicated cereal inhibitors.

- Cation efflux family (Cation\_efflux; Pfam Accession No. PF01545). SEQ ID NO: 321 encodes a polypeptide having sequence similarity to members of the cation efflux family of proteins. Members of this family are integral membrane proteins, that are found to increase tolerance to divalent metal ions such as cadmium, zinc, and cobalt. These proteins are thought to be efflux pumps that remove these ions from cells (Xiong and Jayaswal, *J. Bacteriol.* 180: 4024-4029(1998); Kunito *et al.*, *Biosci. Biotechnol. Biochem.* 60: 699-704(1996)).

DC1 domain (DC1; Pfam Accession No. PF03107). SEQ ID NO: 89 corresponds to a polypeptide having sequence similarity to a DC1 domain. This short domain is rich in cysteines and histidines. The pattern of conservation is similar to that found in DAG\_PE-bind (Pfam Accession No. PF00130), therefore this domain has been termed DC1 for divergent C1 domain. Like the DAG\_PE-bind domain, this domain probably also binds to two zinc ions. The function of proteins with this domain is uncertain, however this domain may bind to molecules such as diacylglycerol. This family are found in plant proteins.

Pneumovirus attachment glycoprotein G (Glycoprotein G; Pfam Accession No. PF00802). SEQ ID NO:995 represents a polypeptide having sequence similarity to members of the Pneumovirus attachment glycoprotein G protein family. This family includes attachment proteins from respiratory syncytial virus. Glycoprotein G has not been shown to have any neuraminidase or hemagglutinin activity. The amino terminus is thought to be cytoplasmic, and the carboxyl terminus extracellular. The extracellular region contains four completely conserved cysteine residues.

NADH-Ubiquinone/plastoquinone (complex I), various chains (oxidored\_q1; Pfam Accession No. PF00361). SEQ ID NO:413 represents a polypeptide having sequence similarity to NADH-Ubiquinone/plastoquinone (complex I), various chains protein family. This family is part of the NADH:ubiquinone oxidoreductase (complex I) which catalyses the transfer of two electrons from NADH to ubiquinone in a reaction that is associated with proton translocation across the membrane (Walker, *Q. Rev. Biophys.* 25: 253-324(1992)). Sub-families within this protein family include NADH-ubiquinone oxidoreductase chain 5; NADH-ubiquinone oxidoreductase chain 2; NADH-ubiquinone oxidoreductase chain 4; and Multicomponent K<sup>+</sup>-H<sup>+</sup>-antiporter.

Protamine P1 (protamine P1; Pfam Accession No. PF00260). SEQ ID NOS:645 and 1217 represent polypeptides having sequence similarity to Protamine P1 protein family. Protamines are small, highly basic proteins, that substitute for histones in sperm chromatin during the haploid phase of spermatogenesis. They pack sperm DNA into a highly condensed, stable and inactive complex. There are two different types of mammalian protamine, called P1 and P2. P1 has been found in all species studied, while P2 is sometimes absent. There also seems to be a single type of avian protamine whose sequence is closely related to that of mammalian P1 (Oliva *et al.*, *J. Biol. Chem.* 264:17627-17630(1989)). A conserved region at the N-terminal extremity of the sequence is used as a signature pattern for this family of proteins. The consensus pattern is: [AV]-R-[NFY]-R-x(2,3)-[ST]-x-S-x-S.

Squash family serine protease inhibitor (squash; Pfam Accession No. PF00299). SEQ ID NO:995 represents a polypeptide having sequence similarity to Squash family serine protease inhibitor proteins. The squash inhibitors form one of a number of serine protease inhibitor families. The proteins, found in the seeds of cucurbitaceae plants (squash, cucumber, balsam pear, etc.), are approximately 30 residues in length, and contain 6 Cys residues, which form 3 disulfide bonds (Bode

*et al., FEBS Lett.* 242: 285-292(1989)). The inhibitors function by being taken up by a serine protease (such as trypsin), which cleaves the peptide bond between Arg/Lys and Ile residues in the N-terminal portion of the protein (Bode *et al., FEBS Lett.* 242: 285-292(1989); Krishnamoorthi *et al., Biochemistry* 31: 898-904(1992)). Structural studies have shown that the inhibitor has an ellipsoidal shape, and is largely composed of beta-turns (Bode *et al., FEBS Lett.* 242: 285-292(1989)). The fold and Cys connectivity of the proteins resembles that of potato carboxypeptidase A inhibitor (Krishnamoorthi *et al., Biochemistry* 31: 898-904(1992)). The pattern used to detect this family of proteins spans the major part of the sequence and includes five of the six cysteines involved in disulfide bonds. The consensus pattern is: C-P-x(5)-C-x(2)-[DN]-x-D-C-x(3)-C-x-C [The five C's are involved in disulfide bonds]

Metallothionein family 5 (Metallothio\_5; Pfam Accession No. PF02067). SEQ ID NO:995 represents a polypeptide having sequence similarity to metallothionein family 5 proteins. Metallothioneins (MT) are small proteins that bind heavy metals, such as zinc, copper, cadmium, and nickel. They have a high content of cysteine residues that bind the metal ions through clusters of thiolate bonds (Kagi, *Meth. Enzymol.* 205: 613-626(1991); Kagi and Kojima, *Experientia Suppl.* 52: 25-61(1987); Kagi and Schaffer, *Biochemistry* 27: 8509-8515(1988)).

Due to limitations in the original classification system of MTs, which did not allow clear differentiation of patterns of structural similarities, either between or within classes, all class I and class II MTs (the proteinaceous sequences) have now been grouped into families of phylogenetically-related and thus alignable sequences. Diptera (*Drosophila*, family 5) MTs are 40-43 residue proteins that contain 10 conserved cysteines arranged in five Cys-X-Cys groups. In particular, the consensus pattern C-G-x(2)-C-x-C-x(2)-Q-x(5)-C-x-C-x(2)-D-C-x-C has been found to be diagnostic of family 5 MTs. The protein is found primarily in the alimentary canal, and its induction is stimulated by ingestion of cadmium or copper (Lastowski *et al., J. Biol. Chem.* 260: 1527-1530(1985)). Mercury, silver and zinc induce the protein to a lesser extent.

Caenorhabditis elegans Sre G protein-coupled chemoreceptor (Sre; Pfam Accession No. PF03125). SEQ ID NO:591 represents a polypeptide having sequence similarity to C. elegans Sre G protein-coupled chemoreceptor family proteins. C. elegans Sre proteins are candidate chemosensory receptors. There are four main recognized groups of such receptors: Odr-10, Sra, Sro, and Srg. Sre (this family), Sra Sra and Srb Srb comprise the Sra group. All of the above receptors are thought to be G protein-coupled seven transmembrane domain proteins (Troemel, *Bioessays* 21:1011-1020 (1999); Troemel *et al., Cell* 83:207-218 (1995)).

Syndecan domain (Syndecan; Pfam Accession No. PF01034). SEQ ID NO:995 corresponds to a polypeptide having a syndecan domain. Syndecans (Bernfield *et al., Annu. Rev. Cell Biol.* 8:365-393(1992); David, *FASEB J.* 7:1023-1030(1993)) are a family of transmembrane heparan sulfate proteoglycans which are implicated in the binding of extracellular matrix components and growth

factors. Syndecans bind a variety of molecules via their heparan sulfate chains and can act as receptors or as co-receptors. Structurally, these proteins consist of four separate domains: a) a signal sequence; b) an extracellular domain (ectodomain) of variable length containing the sites of attachment of the heparan sulfate glycosaminoglycan side chains and whose sequence is not evolutionarily conserved in the various forms of syndecans; c) a transmembrane region; and d) a highly conserved cytoplasmic domain of about 30 to 35 residues which could interact with cytoskeletal proteins.

The signature pattern for syndecans starts with the last residue of the transmembrane region and includes the first 10 residues of the cytoplasmic domain. This region, which contains four basic residues, may act as a stop transfer site. The consensus pattern is: [FY]-R-[IM]-[KR]-K(2)-D-E-G-S-Y.

L1 transposable element (Transposase 22; Pfam Accession No. PF02994). SEQ ID NO:774 represents a polypeptide having an L1 transposable element. Many human L1 elements are capable of retrotransposition and some of these have been shown to exhibit reverse transcriptase (RT) activity (Sassaman *et al.*, *Nat Genet* 16(1):37-43(1997)) although the function of many are, as yet, unknown. There are estimated to be 30-60 active L1 elements reside in the average diploid genome.

WW domain (WW; Pfam Accession No. PF00397). SEQ ID NO:431 represents a polypeptide having WW domain. The WW domain (also known as rsp5 or WWP) is a short conserved region in a number of unrelated proteins, among them dystrophin, responsible for Duchenne muscular dystrophy. This short domain may be repeated up to four times in some proteins (Bork and Sudol, *Trends Biochem. Sci.* 19: 531-533(1994); Andre and Springael, *Biochem. Biophys. Res. Commun.* 205: 1201-1205(1994); Hofmann and Bucher, *FEBS Lett.* 358: 153-157(1995); Sudol *et al.*, *FEBS Lett.* 369: 67-71(1995)). The WW domain binds to proteins with particular proline-motifs, [AP]-P-P-[AP]-Y, and having four conserved aromatic positions that are generally Trp (Chen and Sudol, *Proc. Natl. Acad. Sci. U.S.A.* 92: 7819-7823(1995)). The name WW or WWP derives from the presence of these Trp as well as that of a conserved Pro. The WW domain is frequently associated with other domains typical for proteins in signal transduction processes.

A large variety of proteins containing the WW domain are known. These include; dystrophin, a multidomain cytoskeletal protein; utrophin, a dystrophin-like protein of unknown function; vertebrate YAP protein, substrate of an unknown serine kinase; mouse NEDD-4, involved in the embryonic development and differentiation of the central nervous system; yeast RSP5, similar to NEDD-4 in its molecular organization; rat FE65, a transcription-factor activator expressed preferentially in liver; tobacco DB10 protein and others. The consensus pattern is: W-x(9,11)-[VFY]-[FYW]-x(6,7)-[GSTNE]-[GSTQCR]-[FYW]-x(2)-P.

Example 6: Detection of Differential Expression Using Arrays and source of patient tissue samples



mRNA isolated from samples of cancerous and normal breast and colon tissue obtained from patients were analyzed to identify genes differentially expressed in cancerous and normal cells. Normal and cancerous tissues were collected from patients using laser capture microdissection (LCM) techniques, which techniques are well known in the art (see, e.g., Ohyama *et al.* (2000) *Biotechniques* 29:530-6; Curran *et al.* (2000) *Mol. Pathol.* 53:64-8; Suarez-Quian *et al.* (1999) *Biotechniques* 26:328-35; Simone *et al.* (1998) *Trends Genet* 14:272-6; Conia *et al.* (1997) *J. Clin. Lab. Anal.* 11:28-38; Emmert-Buck *et al.* (1996) *Science* 274:998-1001).

Table 11 (inserted prior to claims) provides information about each patient from which colon tissue samples were isolated, including: the Patient ID ("PT ID") and Path ReportID ("Path ID"), which are numbers assigned to the patient and the pathology reports for identification purposes; the group ("Grp") to which the patients have been assigned; the anatomical location of the tumor ("Anatom Loc"); the primary tumor size ("Size"); the primary tumor grade ("Grade"); the identification of the histopathological grade ("Histo Grade"); a description of local sites to which the tumor had invaded ("Local Invasion"); the presence of lymph node metastases ("Lymph Met"); the incidence of lymph node metastases (provided as a number of lymph nodes positive for metastasis over the number of lymph nodes examined) ("Lymph Met Incid"); the regional lymphnode grade ("Reg Lymph Grade"); the identification or detection of metastases to sites distant to the tumor and their location ("Dist Met & Loc"); the grade of distant metastasis ("Dist Met Grade"); and general comments about the patient or the tumor ("Comments"). Histopathology of all primary tumors indicated the tumor was adenocarcinoma except for Patient ID Nos. 130 (for which no information was provided), 392 (in which greater than 50% of the cells were mucinous carcinoma), and 784 (adenosquamous carcinoma). Extranodal extensions were described in three patients, Patient ID Nos. 784, 789, and 791. Lymphovascular invasion was described in Patient ID Nos. 128, 278, 517, 534, 784, 786, 789, 791, 890, and 892. Crohn's-like infiltrates were described in seven patients, Patient ID Nos. 52, 264, 268, 392, 393, 784, and 791.

Table 12 (below) provides information about each patient from which the breast tissue samples were isolated, including: 1) the "Pat Num", a number assigned to the patient for identification purposes; 2) the "Histology", which indicates whether the tumor was characterized as an intraductal carcinoma (IDC) or ductal carcinoma in situ (DCIS); 3) the incidence of lymph node metastases (LMF), represented as the number of lymph nodes positive to metastases out of the total number examined in the patient; 4) the "Tumor Size"; 5) "TNM Stage", which provides the tumor grade (T#), where the number indicates the grade and "p" indicates that the tumor grade is a pathological classification; regional lymph node metastasis (N#), where "0" indicates no lymph node metastases were found, "1" indicates lymph node metastases were found, and "X" means information not available; and the identification or detection of metastases to sites distant to the tumor and their

location (M#), with "X" indicating that no distant metastases were reported; and the stage of the tumor ("Stage Grouping"). "nr" indicates "no reported".

**Table 12.** Breast cancer patient data.

Pat Num	Histology	LMF	Tumor Size	TNM Stage	Stage Grouping
280	IDC, DCIS+D2	nr	2 cm	T2NXMX	probable Stage II
284	IDC, DCIS	0/16	2 cm	T2pN0MX	Stage II
285	IDC, DCIS	nr	4.5 cm	T2NXMX	probable Stage II
291	IDC, DCIS	0/24	4.5 cm	T2pN0MX	Stage II
302	IDC, DCIS	nr	2.2 cm	T2NXMX	probable Stage II
375	IDC, DCIS	nr	1.5 cm	T1NXMX	probable Stage I
408	IDC	0/23	3.0 cm	T2pN0MX	Stage II
416	IDC	0/6	3.3 cm	T2pN0MX	Stage II
421	IDC, DCIS	nr	3.5 cm	T2NXMX	probable Stage II
459	IDC	2/5	4.9 cm	T2pN1MX	Stage II
465	IDC	0/10	6.5 cm	T3pN0MX	Stage II
470	IDC, DCIS	0/6	2.5 cm	T2pN0MX	Stage II
472	IDC, DCIS	6/45	5.0+ cm	T3pN1MX	Stage III
474	IDC	0/18	6.0 cm	T3pN0MX	Stage II
476	IDC	0/16	3.4 cm	T2pN0MX	Stage II
605	IDC, DCIS	1/25	5.0 cm	T2pN1MX	Stage II
649	IDC, DCIS	1/29	4.5 cm	T2pN1MX	Stage II

Identification of differentially expressed genes

- 5 cDNA probes were prepared from total RNA isolated from the patient cells described above. Since LCM provides for the isolation of specific cell types to provide a substantially homogenous cell sample, this provided for a similarly pure RNA sample.

- Total RNA was first reverse transcribed into cDNA using a primer containing a T7 RNA polymerase promoter, followed by second strand DNA synthesis. cDNA was then transcribed *in vitro* to produce antisense RNA using the T7 promoter-mediated expression (see, e.g., Luo *et al.* (1999) *Nature Med* 5:117-122), and the antisense RNA was then converted into cDNA. The second set of cDNAs were again transcribed *in vitro*, using the T7 promoter, to provide antisense RNA. Optionally, the RNA was again converted into cDNA, allowing for up to a third round of T7-mediated amplification to produce more antisense RNA. Thus the procedure provided for two or three rounds of *in vitro* transcription to produce the final RNA used for fluorescent labeling.

Fluorescent probes were generated by first adding control RNA to the antisense RNA mix, and producing fluorescently labeled cDNA from the RNA starting material. Fluorescently labeled cDNAs prepared from the tumor RNA sample were compared to fluorescently labeled cDNAs prepared from normal cell RNA sample. For example, the cDNA probes from the normal cells were

labeled with Cy3 fluorescent dye (green) and the cDNA probes prepared from the tumor cells were labeled with Cy5 fluorescent dye (red), and vice versa.

Each array used had an identical spatial layout and control spot set. Each microarray was divided into two areas, each area having an array with, on each half, twelve groupings of 32 x 12 spots, for a total of about 9,216 spots on each array. The two areas are spotted identically which provide for at least two duplicates of each clone per array.

Polynucleotides for use on the arrays were obtained from both publicly available sources and from cDNA libraries generated from selected cell lines and patient tissues. PCR products of from about 0.5kb to 2.0 kb amplified from these sources were spotted onto the array using a Molecular Dynamics Gen III spotter according to the manufacturer's recommendations. The first row of each of the 24 regions on the array had about 32 control spots, including 4 negative control spots and 8 test polynucleotides. The test polynucleotides were spiked into each sample before the labeling reaction with a range of concentrations from 2-600 pg/slide and ratios of 1:1. For each array design, two slides were hybridized with the test samples reverse-labeled in the labeling reaction. This provided for about four duplicate measurements for each clone, two of one color and two of the other, for each sample.

The differential expression assay was performed by mixing equal amounts of probes from tumor cells and normal cells of the same patient ("matched") or from tumor cells and normal cells of different patients ("unmatched") (i.e., the tumor cells are from one patient and the normal cells are from a different patient). The arrays were prehybridized by incubation for about 2 hrs at 60°C in 5X SSC/0.2% SDS/1 mM EDTA, and then washed three times in water and twice in isopropanol. Following prehybridization of the array, the probe mixture was then hybridized to the array under conditions of high stringency (overnight at 42°C in 50% formamide, 5X SSC, and 0.2% SDS. After hybridization, the array was washed at 55°C three times as follows: 1) first wash in 1X SSC/0.2% SDS; 2) second wash in 0.1X SSC/0.2% SDS; and 3) third wash in 0.1X SSC.

The arrays were then scanned for green and red fluorescence using a Molecular Dynamics Generation III dual color laser-scanner/detector. The images were processed using BioDiscovery Autogene software, and the data from each scan set normalized to provide for a ratio of expression relative to normal. Data from the microarray experiments was analyzed according to the algorithms described in U.S. application serial no. 60/252,358, filed November 20, 2000, by E.J. Moler, M.A. Boyle, and F.M. Randazzo, and entitled "Precision and accuracy in cDNA microarray data," which application is specifically incorporated herein by reference.

The experiment was repeated, this time labeling the two probes with the opposite color in order to perform the assay in both "color directions." Each experiment was sometimes repeated with two more slides (one in each color direction). The level fluorescence for each sequence on the array expressed as a ratio of the geometric mean of 8 replicate spots/genes from the four arrays or 4 replicate spots/gene from 2 arrays or some other permutation. The data were normalized using the spiked

positive controls present in each duplicated area, and the precision of this normalization was included in the final determination of the significance of each differential. The fluorescent intensity of each spot was also compared to the negative controls in each duplicated area to determine which spots have detected significant expression levels in each sample.

5 A statistical analysis of the fluorescent intensities was applied to each set of duplicate spots to assess the precision and significance of each differential measurement, resulting in a p-value testing the null hypothesis that there is no differential in the expression level between the tumor and normal samples of each patient in matched samples or between tumor and normal samples of tissue from different patients in unmatched samples. During initial analysis of the microarrays, the hypothesis was  
10 accepted if  $p > 10^{-3}$ , and the differential ratio was set to 1.000 for those spots. All other spots have a significant difference in expression between the tumor and normal sample. If the tumor sample has detectable expression and the normal does not, the ratio is truncated at 1000 since the value for expression in the normal sample would be zero, and the ratio would not be a mathematically useful value (e.g., infinity). If the normal sample has detectable expression and the tumor does not, the ratio  
15 is truncated to 0.001, since the value for expression in the tumor sample would be zero and the ratio would not be a mathematically useful value. These latter two situations are referred to herein as "on/off." Database tables were populated using a 95% confidence level ( $p > 0.05$ ).

Table 13 (inserted prior to claims) provides the results for gene products expressed by at least 2-fold or greater in cancerous prostate, colon, or breast tissue samples relative to normal tissue  
20 samples in at least 20% of the patients tested. Table 13 includes: 1) the SEQ ID NO ("SEQ ID") assigned to each sequence for use in the present specification; 2) the sequence name ("SEQ NAME") used as an internal identifier of the sequence; 3) the name assigned to the clone from which the sequence was isolated ("CLONE ID"); 4) the percentage of patients tested in which expression levels (e.g., as message level) of the gene was at least 2-fold greater in cancerous breast tissue than in  
25 matched normal tissue ("BREAST PATIENTS  $\geq 2x$ "); 5) the breast number ratios, indicating the number of patients upon which the provided ratio using matched breast tissue was based ("BREAST NUM RATIOS"); 6) the percentage of patients tested in which expression levels (e.g., as message level) of the gene was at least 2-fold greater in cancerous colon tissue than in matched normal tissue ("COLON PATIENTS  $\geq 2x$ "); 7) the colon number ratios, indicating the number of patients upon  
30 which the provided ratio using matched colon tissue was based ("COLON NUM RATIOS"); 8) the percentage of patients tested in which expression levels (e.g., as message level) of the gene was at least 2-fold greater in cancerous colon tissue than in unmatched normal tissue ("COLON UM  $\geq 2x$ "); 9) the unmatched colon number ratios, indicating the number of patients upon which the provided ratio using unmatched colon tissue was based ("COLON UM NUM RATIOS").

35 Table 16 (inserted prior to claims) provides the results for other gene products expressed by at least 2-fold or greater in cancerous prostate, colon, or breast tissue samples, which may be

metastasized cancer samples, relative to normal tissue samples in at least 20% of the patients tested. For each set of data (i.e., the percentage of patients in which a particular sequence is up-regulated in a cancer tissue) the number of patients (Colon Cancer Patients; Colon Unmatched Met Patients and Colon Match Met Patients) is shown. If a sample is matched, it is matched to a sample from the same patient, if a sample is unmatched, the results obtained from that sample are compared to a pooled sample of an appropriate tissue type from the patients. If a sample is not from a metastasized tissue, it is from a primary tumor.

These data provide evidence that the genes represented by the polynucleotides having the indicated sequences are differentially expressed in breast, prostate, cancer as compared to normal non-cancerous breast tissue and are differentially expressed in colon cancer as compared to normal non-cancerous colon tissue

The above methods can be performed to identify genes differentially expressed in cancerous and normal cells of any type of tissue, such as prostate, lung, colon, breast, and the like.

#### Example 7: Antisense Regulation of Gene Expression

The expression of the differentially expressed genes represented by the polynucleotides in the cancerous cells can be further analyzed using antisense knockout technology to confirm the role and function of the gene product in tumorigenesis, *e.g.*, in promoting a metastatic phenotype.

Methods for analysis using antisense technology are well known in the art. For example, a number of different oligonucleotides complementary to the mRNA generated by the differentially expressed genes identified herein can be designed as antisense oligonucleotides, and tested for their ability to suppress expression of the genes. Sets of antisense oligomers specific to each candidate target are designed using the sequences of the polynucleotides corresponding to a differentially expressed gene and the software program HYBsimulator Version 4 (available for Windows 95/Windows NT or for Power Macintosh, RNAture, Inc. 1003 Health Sciences Road, West, Irvine, CA 92612 USA). Factors considered when designing antisense oligonucleotides include: 1) the expression of the differentially expressed genes represented by the polynucleotides in the cancerous cells can be analyzed using antisense knockout technology to confirm the role and function of the gene product in tumorigenesis, *e.g.*, in promoting a metastatic phenotype.

A number of different oligonucleotides complementary to the mRNA generated by the differentially expressed genes identified herein can be designed as potential antisense oligonucleotides, and tested for their ability to suppress expression of the genes. Sets of antisense oligomers specific to each candidate target are designed using the sequences of the polynucleotides corresponding to a differentially expressed gene and the software program HYBsimulator Version 4 (available for Windows 95/Windows NT or for Power Macintosh, RNAture, Inc. 1003 Health Sciences Road, West, Irvine, CA 92612 USA). Factors that are considered when designing antisense oligonucleotides include: 1) the secondary structure of oligonucleotides; 2) the secondary structure of

the target gene; 3) the specificity with no or minimum cross-hybridization to other expressed genes; 4) stability; 5) length and 6) terminal GC content. The antisense oligonucleotide is designed so that it will hybridize to its target sequence under conditions of high stringency at physiological temperatures (e.g., an optimal temperature for the cells in culture to provide for hybridization in the cell, e.g., about 37°C), but with minimal formation of homodimers.

Using the sets of oligomers and the HYBSimulator program, three to ten antisense oligonucleotides and their reverse controls are designed and synthesized for each candidate mRNA transcript, which transcript is obtained from the gene corresponding to the target polynucleotide sequence of interest. Once synthesized and quantitated, the oligomers are screened for efficiency of a transcript knock-out in a panel of cancer cell lines. The efficiency of the knock-out is determined by analyzing mRNA levels using lightcycler quantification. The oligomers that resulted in the highest level of transcript knock-out, wherein the level was at least about 50%, preferably about 80-90%, up to 95% or more up to undetectable message, are selected for use in a cell-based proliferation assay, an anchorage independent growth assay, and an apoptosis assay.

The ability of each designed antisense oligonucleotide to inhibit gene expression is tested through transfection into LNCaP, PC3, 22Rv1, MDA-PCA-2b, or DU145 prostate carcinoma cells. For each transfection mixture, a carrier molecule (such as a lipid, lipid derivative, lipid-like molecule, cholesterol, cholesterol derivative, or cholesterol-like molecule) is prepared to a working concentration of 0.5 mM in water, sonicated to yield a uniform solution, and filtered through a 0.45 µm PVDF membrane. The antisense or control oligonucleotide is then prepared to a working concentration of 100 µM in sterile Millipore water. The oligonucleotide is further diluted in OptiMEM™ (Gibco/BRL), in a microfuge tube, to 2 µM, or approximately 20 µg oligo/ml of OptiMEM™. In a separate microfuge tube, the carrier molecule, typically in the amount of about 1.5-2 nmol carrier/µg antisense oligonucleotide, is diluted into the same volume of OptiMEM™ used to dilute the oligonucleotide. The diluted antisense oligonucleotide is immediately added to the diluted carrier and mixed by pipetting up and down. Oligonucleotide is added to the cells to a final concentration of 30 nM.

The level of target mRNA that corresponds to a target gene of interest in the transfected cells is quantitated in the cancer cell lines using the Roche LightCycler™ real-time PCR machine. Values for the target mRNA are normalized versus an internal control (e.g., beta-actin). For each 20 µl reaction, extracted RNA (generally 0.2-1 µg total) is placed into a sterile 0.5 or 1.5 ml microcentrifuge tube, and water is added to a total volume of 12.5 µl. To each tube is added 7.5 µl of a buffer/enzyme mixture, prepared by mixing (in the order listed) 2.5 µl H<sub>2</sub>O, 2.0 µl 10X reaction buffer, 10 µl oligo dT (20 pmol), 1.0 µl dNTP mix (10 mM each), 0.5 µl RNAsin® (20u) (Ambion, Inc., Hialeah, FL), and 0.5 µl MMLV reverse transcriptase (50u) (Ambion, Inc.). The contents are mixed by pipetting up

and down, and the reaction mixture is incubated at 42°C for 1 hour. The contents of each tube are centrifuged prior to amplification.

An amplification mixture is prepared by mixing in the following order: 1X PCR buffer II, 3 mM MgCl<sub>2</sub>, 140 μM each dNTP, 0.175 pmol each oligo, 1:50,000 dil of SYBR® Green, 0.25 mg/ml BSA, 1 unit *Taq* polymerase, and H<sub>2</sub>O to 20 μl. (PCR buffer II is available in 10X concentration from Perkin-Elmer, Norwalk, CT). In 1X concentration it contains 10 mM Tris pH 8.3 and 50 mM KCl. SYBR® Green (Molecular Probes, Eugene, OR) is a dye which fluoresces when bound to double stranded DNA. As double stranded PCR product is produced during amplification, the fluorescence from SYBR® Green increases. To each 20 μl aliquot of amplification mixture, 2 μl of template RT is added, and amplification is carried out according to standard protocols. The results are expressed as the percent decrease in expression of the corresponding gene product relative to non-transfected cells, vehicle-only transfected (mock-transfected) cells, or cells transfected with reverse control oligonucleotides.

#### Example 8: Effect of Expression on Proliferation

The effect of gene expression on the inhibition of cell proliferation can be assessed in metastatic breast cancer cell lines (MDA-MB-231 ("231")); SW620 colon colorectal carcinoma cells; SKOV3 cells (a human ovarian carcinoma cell line); or LNCaP, PC3, 22Rv1, MDA-PCA-2b, or DU145 prostate cancer cells.

Cells are plated to approximately 60-80% confluency in 96-well dishes. Antisense or reverse control oligonucleotide is diluted to 2 μM in OptiMEM™. The oligonucleotide-OptiMEM™ can then be added to a delivery vehicle, which delivery vehicle can be selected so as to be optimized for the particular cell type to be used in the assay. The oligo/delivery vehicle mixture is then further diluted into medium with serum on the cells. The final concentration of oligonucleotide for all experiments can be about 300 nM.

Antisense oligonucleotides are prepared as described above (see Example 3). Cells are transfected overnight at 37°C and the transfection mixture is replaced with fresh medium the next morning. Transfection is carried out as described above in Example 8.

Those antisense oligonucleotides that result in inhibition of proliferation of SW620 cells indicate that the corresponding gene plays a role in production or maintenance of the cancerous phenotype in cancerous colon cells. Those antisense oligonucleotides that inhibit proliferation in SKOV3 cells represent genes that play a role in production or maintenance of the cancerous phenotype in cancerous breast cells. Those antisense oligonucleotides that result in inhibition of proliferation of MDA-MB-231 cells indicate that the corresponding gene plays a role in production or maintenance of the cancerous phenotype in cancerous ovarian cells. Those antisense oligonucleotides that inhibit proliferation in LNCaP, PC3, 22Rv1, MDA-PCA-2b, or DU145 cells represent genes that play a role in production or maintenance of the cancerous phenotype in cancerous prostate cells.

Using the following antisense oligonucleotides: TTGGTTCCTCCCAAGACAAGCCGTGAC (SEQ ID NO:1543); TCTCAACGCTACCAGGCACTCCCTTG (SEQ ID NO:1544); GCACAGCCCAAGTCAAAGGCATTA (SEQ ID NO:1545); CAGGCACTCCTTGGTCAAATGTGGG (SEQ ID NO:1546);

5 GGACAGGGAAGGAGAGGCTAGTCA (SEQ ID NO:1547) and TGCATTCTCTCCACATCTCAACGC (SEQ ID NO:1548, corresponding to a glutathione transferase omega identified by SEQ ID NOS: 1377 and 1541 (Chiron Candidate Id 21), were used to inhibit proliferation of SW620 colon colorectal carcinoma cells. These antisense molecules reduced glutathione transferase omega RNA expression by approximately 90%.

10 Example 9: Effect of Gene Expression on Cell Migration

The effect of gene expression on the inhibition of cell migration can be assessed in LNCaP, PC3, 22Rv1, MDA-PCA-2b, or DU145 prostate cancer cells using static endothelial cell binding assays, non-static endothelial cell binding assays, and transmigration assays.

For the static endothelial cell binding assay, antisense oligonucleotides are prepared as

15 described above (see Example 8). Two days prior to use, prostate cancer cells (CaP) are plated and transfected with antisense oligonucleotide as described above (see Examples 3 and 4). On the day before use, the medium is replaced with fresh medium, and on the day of use, the medium is replaced with fresh medium containing 2  $\mu$ M CellTracker green CMFDA (Molecular Probes, Inc.) and cells are incubated for 30 min. Following incubation, CaP medium is replaced with fresh medium (no

20 CMFDA) and cells are incubated for an additional 30-60 min. CaP cells are detached using CMF PBS/2.5 mM EDTA or trypsin, spun and resuspended in DMEM/1% BSA/ 10 mM HEPES pH 7.0. Finally, CaP cells are counted and resuspended at a concentration of  $1 \times 10^6$  cells/ml.

Endothelial cells (EC) are plated onto 96-well plates at 40-50% confluence 3 days prior to use. On the day of use, EC are washed 1X with PBS and 50 $\lambda$  DMDM/1%BSA/10mM HEPES pH 7

25 is added to each well. To each well is then added 50K (50 $\lambda$ ) CaP cells in DMEM/1% BSA/ 10mM HEPES pH 7. The plates are incubated for an additional 30 min and washed 5X with PBS containing Ca<sup>++</sup> and Mg<sup>++</sup>. After the final wash, 100  $\mu$ L PBS is added to each well and fluorescence is read on a fluorescent plate reader (Ab492/Em 516 nm).

For the non-static endothelial cell binding assay, CaP are prepared as described above. EC

30 are plated onto 24-well plates at 30-40% confluence 3 days prior to use. On the day of use, a subset of EC are treated with cytokine for 6 hours then washed 2X with PBS. To each well is then added 150-200K CaP cells in DMEM/1% BSA/ 10mM HEPES pH 7. Plates are placed on a rotating shaker (70 RPM) for 30 min and then washed 3X with PBS containing Ca<sup>++</sup> and Mg<sup>++</sup>. After the final wash, 500  $\mu$ L PBS is added to each well and fluorescence is read on a fluorescent plate reader (Ab492/Em 516

35 nm).



For the transmigration assay, CaP are prepared as described above with the following changes. On the day of use, CaP medium is replaced with fresh medium containing 5  $\mu$ M CellTracker green CMFDA (Molecular Probes, Inc.) and cells are incubated for 30 min. Following incubation, CaP medium is replaced with fresh medium (no CMFDA) and cells are incubated for an additional 30-60 min. CaP cells are detached using CMF PBS/2.5 mM EDTA or trypsin, spun and resuspended in EGM-2-MV medium. Finally, CaP cells are counted and resuspended at a concentration of  $1 \times 10^6$  cells/ml.

EC are plated onto FluorBlok transwells (BD Biosciences) at 30-40% confluence 5-7 days before use. Medium is replaced with fresh medium 3 days before use and on the day of use. To each transwell is then added 50K labeled CaP. 30 min prior to the first fluorescence reading, 10  $\mu$ g of FITC-dextran (10K MW) is added to the EC plated filter. Fluorescence is then read at multiple time points on a fluorescent plate reader (Ab492/Em 516 nm).

Those antisense oligonucleotides that result in inhibition of binding of LNCaP, PC3, 22Rv1, MDA-PCA-2b, or DU145 prostate cancer cells to endothelial cells indicate that the corresponding gene plays a role in the production or maintenance of the cancerous phenotype in cancerous prostate cells. Those antisense oligonucleotides that result in inhibition of endothelial cell transmigration by LNCaP, PC3, 22Rv1, MDA-PCA-2b, or DU145 prostate cancer cells indicate that the corresponding gene plays a role in the production or maintenance of the cancerous phenotype in cancerous prostate cells.

#### Example 10: Effect of Gene Expression on Colony Formation

The effect of gene expression upon colony formation of SW620 cells, SKOV3 cells, MD-MBA-231 cells, LNCaP cells, PC3 cells, 22Rv1 cells, MDA-PCA-2b cells, and DU145 cells can be tested in a soft agar assay. Soft agar assays are conducted by first establishing a bottom layer of 2 ml of 0.6% agar in media plated fresh within a few hours of layering on the cells. The cell layer is formed on the bottom layer by removing cells transfected as described above from plates using 0.05% trypsin and washing twice in media. The cells are counted in a Coulter counter, and resuspended to  $10^6$  per ml in media. 10  $\mu$ l aliquots are placed with media in 96-well plates (to check counting with WST1), or diluted further for the soft agar assay. 2000 cells are plated in 800  $\mu$ l 0.4% agar in duplicate wells above 0.6% agar bottom layer. After the cell layer agar solidifies, 2 ml of media is dribbled on top and antisense or reverse control oligo (produced as described in Example 8) is added without delivery vehicles. Fresh media and oligos are added every 3-4 days. Colonies form in 10 days to 3 weeks. Fields of colonies are counted by eye. Wst-1 metabolism values can be used to compensate for small differences in starting cell number. Larger fields can be scanned for visual record of differences.

Those antisense oligonucleotides that result in inhibition of colony formation of SW620 cells indicate that the corresponding gene plays a role in production or maintenance of the cancerous

phenotype in cancerous colon cells. Those antisense oligonucleotides that inhibit colony formation in SKOV3 cells represent genes that play a role in production or maintenance of the cancerous phenotype in cancerous breast cells. Those antisense oligonucleotides that result in inhibition of colony formation of MDA-MB-231 cells indicate that the corresponding gene plays a role in production or maintenance of the cancerous phenotype in cancerous ovarian cells. Those antisense oligonucleotides that inhibit colony formation in LNCaP, PC3, 22Rv1, MDA-PCA-2b, or DU145 cells represent genes that play a role in production or maintenance of the cancerous phenotype in cancerous prostate cells.

Example 11: Induction of Cell Death upon Depletion of Polypeptides by Depletion of mRNA

("Antisense Knockout")

In order to assess the effect of depletion of a target message upon cell death, LNCaP, PC3, 22Rv1, MDA-PCA-2b, or DU145 cells, or other cells derived from a cancer of interest, can be transfected for proliferation assays. For cytotoxic effect in the presence of cisplatin (cis), the same protocol is followed but cells are left in the presence of 2  $\mu$ M drug. Each day, cytotoxicity is monitored by measuring the amount of LDH enzyme released in the medium due to membrane damage. The activity of LDH is measured using the Cytotoxicity Detection Kit from Roche Molecular Biochemicals. The data is provided as a ratio of LDH released in the medium vs. the total LDH present in the well at the same time point and treatment (rLDH/tLDH). A positive control using antisense and reverse control oligonucleotides for BCL2 (a known anti-apoptotic gene) is included; loss of message for BCL2 leads to an increase in cell death compared with treatment with the control oligonucleotide (background cytotoxicity due to transfection).

Example 12: Functional Analysis of Gene Products Differentially Expressed in Cancer

The gene products of sequences of a gene differentially expressed in cancerous cells can be further analyzed to confirm the role and function of the gene product in tumorigenesis, e.g., in promoting or inhibiting development of a metastatic phenotype. For example, the function of gene products corresponding to genes identified herein can be assessed by blocking function of the gene products in the cell. For example, where the gene product is secreted or associated with a cell surface membrane, blocking antibodies can be generated and added to cells to examine the effect upon the cell phenotype in the context of, for example, the transformation of the cell to a cancerous, particularly a metastatic, phenotype. In order to generate antibodies, a clone corresponding to a selected gene product is selected, and a sequence that represents a partial or complete coding sequence is obtained. The resulting clone is expressed, the polypeptide produced isolated, and antibodies generated. The antibodies are then combined with cells and the effect upon tumorigenesis assessed.

Where the gene product of the differentially expressed genes identified herein exhibits sequence homology to a protein of known function (e.g., to a specific kinase or protease) and/or to a protein family of known function (e.g., contains a domain or other consensus sequence present in a

protease family or in a kinase family), then the role of the gene product in tumorigenesis, as well as the activity of the gene product, can be examined using small molecules that inhibit or enhance function of the corresponding protein or protein family.

Additional functional assays include, but are not necessarily limited to, those that analyze the effect of expression of the corresponding gene upon cell cycle and cell migration. Methods for performing such assays are well known in the art.

Example 13: Deposit Information.

Deposits of the biological materials in the tables referenced below were made with either the Agricultural Research Service Culture Collection (NRRL), 1815 North University Street, Peoria, Illinois 61604, or with the American Type Culture Collection (ATCC), 10801 University Blvd., Manassas, VA 20110-2209, under the provisions of the Budapest Treaty, on or before the filing date of the present application. The accession number indicated is assigned after successful viability testing, and the requisite fees were paid. Access to said cultures will be available during pendency of the patent application to one determined by the Commissioner to be entitled to such under 37 C.F.R. §1.14 and 35 U.S.C. §122. All restriction on availability of said cultures to the public will be irrevocably removed upon the granting of a patent based upon the application. Moreover, the designated deposits will be maintained for a period of thirty (30) years from the date of deposit, or for five (5) years after the last request for the deposit; or for the enforceable life of the U.S. patent, whichever is longer. Should a culture become nonviable or be inadvertently destroyed, or, in the case of plasmid-containing strains, lose its plasmid, it will be replaced with a viable culture(s) of the same taxonomic description.

These deposits are provided merely as a convenience to those of skill in the art, and are not an admission that a deposit is required. A license may be required to make, use, or sell the deposited materials, and no such license is hereby granted. The deposit below was received by the ATCC on or before the filing date of the present application.

**Table 14. Cell Lines Deposited with ATCC**

Cell Line	Deposit Date	ATCC Accession No.	CMCC Accession No.
KM12L4-A	March 19, 1998	CRL-12496	11606
Km12C	May 15, 1998	CRL-12533	11611
MDA-MB-231	May 15, 1998	CRL-12532	10583
MCF-7	October 9, 1998	CRL-12584	10377

In addition, pools of selected clones, as well as libraries containing specific clones, were assigned an "ES" number and a "CMCC" number (both internal references) and deposited with the NRRL. Table 15 (inserted before the claims) provides the NRRL Accession Nos. of the clones

deposited as libraries named ES219-ES225 (CMCC5471-CMCC5477, respectively) on November 1, 2001, and of the clones deposited as a library named ES226 (CMCC5478) on November 7, 2001.

Retrieval of Individual Clones from Deposit of Pooled Clones. Where the biological deposit  
5 is composed of a pool of cDNA clones or a library of cDNA clones, the deposit was prepared by first transfecting each of the clones into separate bacterial cells. The clones in the pool or library were then deposited as a pool of equal mixtures in the composite deposit. Particular clones can be obtained from the composite deposit using methods well known in the art. For example, a bacterial cell containing a particular clone can be identified by isolating single colonies, and identifying colonies containing the  
10 specific clone through standard colony hybridization techniques, using an oligonucleotide probe or probes designed to specifically hybridize to a sequence of the clone insert (e.g., a probe based upon unmasked sequence of the encoded polynucleotide having the indicated SEQ ID NO). The probe should be designed to have a  $T_m$  of approximately 80°C (assuming 2°C for each A or T and 4°C for each G or C). Positive colonies can then be picked, grown in culture, and the recombinant clone  
15 isolated. Alternatively, probes designed in this manner can be used to PCR to isolate a nucleic acid molecule from the pooled clones according to methods well known in the art, e.g., by purifying the cDNA from the deposited culture pool, and using the probes in PCR reactions to produce an amplified product having the corresponding desired polynucleotide sequence.

Although the foregoing invention has been described in some detail by way of illustration and  
20 example for purposes of clarity of understanding, it is readily apparent to those of ordinary skill in the art in light of the teachings of this invention that certain changes and modifications may be made thereto without departing from the spirit or scope of the appended claims. Those skilled in the art will recognize, or be able to ascertain, using not more than routine experimentation, many equivalents to the specific embodiments of the invention described herein. Such specific embodiments and  
25 equivalents are intended to be encompassed by the following claims.

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
1	3538.O24.GZ43_504925	AF047717	Streptomyces chrysomallus actinomycin synthetase II (acmB) gene, complete cds	1.17E-04
2	3538.P11.GZ43_504718	AF111848	Homo sapiens PRO0529 mRNA, complete cds	2.00E-06
3	3541.A04.GZ43_504975	X58178	S.pyogenes for enm41 gene	5.00E-06
4	3541.A05.GZ43_504991	AF190638	Mus musculus nephrin NPHS1 (Nphs1) gene, partial cds	2.00E-06
5	3541.A16.GZ43_505167	AB024689	Mus musculus gene, exon 3, partial sequence	6.00E-06
6	3541.A23.GZ43_505279	M14155	Human insulin-like growth factor (IGF-I) IB gene, exon 4	3.00E-06
7	3541.B04.GZ43_504976	U32801	Haemophilus influenzae Rd section 116 of 163 of the complete genome	1.10E-05
8	3541.B17.GZ43_505184	X89398	H.sapiens ung gene for uracil DNA-glycosylase	1.21E-04
9	3538.G08.GZ43_504661	AF270390	Staphylococcus epidermidis strain SR1 clone step.4045d08 genomic sequence	3.00E-06
10	3538.G17.GZ43_504805	AC006623	Caenorhabditis elegans clone C52E2, complete sequence	4.00E-06
11	3538.G19.GZ43_504837	AB042425	Homo sapiens Pim-2h, hUGT2, hUGT1, genes for pim-2 proto-oncogene homolog, UDP-galactose transporter 1, UDP-galactose transporter 2, complete cds	6.60E-11
12	3538.G22.GZ43_504885	L08338	Human immunodeficiency virus type 1 proviral envelope glycoprotein gene V3 region from A196/4537, clone 6 (from adult)	3.10E-07
13	3538.H05.GZ43_504614	AE006731	Sulfolobus solfataricus section 90 of 272 of the complete genome	2.00E-06
14	3538.H21.GZ43_504870	AL121807	S.pombe chromosome III cosmid c132	1.30E-05
15	3538.I08.GZ43_504663	AF186379	Homo sapiens ligand effect modulator-6 (LEM6) mRNA, complete cds	8.00E-10
16	3538.I13.GZ43_504743	AC007658	Arabidopsis thaliana chromosome II section 216 of 255 of the complete sequence. Sequence from clones F2711	3.30E-08
17	3538.J22.GZ43_504888	X04616	Anacystis nidulans R2 psbAI gene for photosystem II Q(B) protein	8.90E-07
18	3538.K12.GZ43_504729	X91656	M.musculus Srp20 gene	4.40E-05
19	3538.K23.GZ43_504905	M62849	Human papillomavirus ORFs	4.40E-07
20	3538.L16.GZ43_504794	AE001382	Plasmodium falciparum chromosome 2, section 19 of 73 of the complete sequence	7.00E-06
21	3538.M02.GZ43_504571	U07976	Human T cell receptor beta (TCRBV7S2, TCRBV13S2-1, TCRBV6S7-1) genes, TCRBV deleted 2 haplotype, partial cds	7.00E-06
22	3538.M05.GZ43_504619	AC079878	Homo sapiens BAC clone RP11-343P21 from 7, complete sequence	1.40E-07
23	3538.M08.GZ43_504667	AF182668	Zenaida galapagoensis beta-fibrinogen gene, partial sequence	4.70E-08

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
24	3538.N20.GZ43_504860	AB033411	Taenia crassiceps mitochondrial gene for cytochrome c oxidase subunit 1, partial cds	6.80E-07
25	3538.O07.GZ43_504653	X68019	Feline Immunodeficiency Virus GAG gene	4.00E-06
26	3541.E11.GZ43_505091	M73447	Human repeat polymorphism at locus D9S59	3.00E-08
27	3541.E14.GZ43_505139	AJ243419	Acanthospora trappei partial 18S rRNA, 5.8S rRNA and partial 28S rRNA genes and internal transcribed spacers 1 and 2 (ITS1, ITS2), isolate AU 219	1.10E-07
28	3541.E15.GZ43_505155	U13679	Human lactate dehydrogenase-A (LDH-A) gene, promoter region	3.50E-10
29	3541.G17.GZ43_505189	AE004851	Pseudomonas aeruginosa PA01, section 412 of 529 of the complete genome	1.30E-05
30	3541.H14.GZ43_505142	AJ252202	Drosophila melanogaster D-COQ7 gene for putative COQ7 isologue, exons 1-3	9.00E-06
31	3541.I15.GZ43_505159	X98371	D.subobscura sex-lethal gene	6.00E-06
32	3541.I17.GZ43_505191	AK023918	Homo sapiens cDNA FLJ13856 fis, clone THYRO1000988	1.70E-22
33	3541.I18.GZ43_505207	AF329081	Bos taurus AMP-activated protein kinase gamma-1 (PRKAG1) gene, partial cds	5.30E-33
34	3541.J19.GZ43_505224	AF002749	Psychotria urceolata ribosomal protein S16 (rps16) gene, chloroplast gene encoding chloroplast protein, partial intron	3.01E-03
35	3541.K09.GZ43_505065	AF027607	Gallus gallus L-type voltage-gated calcium channel alpha1D subunit ChCaChA1D precursor mRNA, complete intron sequence	9.00E-06
36	3541.L19.GZ43_505226	AE003949	Xylella fastidiosa 9a5c, section 95 of 229 of the complete genome	2.00E-06
37	3541.M02.GZ43_504955	BC004556	Homo sapiens, Similar to CG7083 gene product, clone MGC:10534 IMAGE3957147, mRNA, complete cds	6.20E-07
38	3541.M07.GZ43_505035	X05616	Kangaroo rat repetitive DNA with insertion sequence	4.80E-08
39	3541.M18.GZ43_505211	M81888	Parvovirus LuII DNA sequence	6.60E-05
40	3541.O04.GZ43_504989	AF081828	Ixodes hexagonus mitochondrial DNA, complete genome	3.00E-06
41	3541.O13.GZ43_505133	AK026465	Homo sapiens cDNA: FLJ22812 fis, clone KAI2955	8.00E-06
42	3541.O23.GZ43_505293	X54859	Porcine TNF-alpha and TNF-beta genes for tumour necrosis factors alpha and beta, respectively	2.90E-05
43	3541.P05.GZ43_505006	AE006642	Sulfolobus solfataricus section 1 of 272 of the complete genome	3.50E-05
44	3541.P22.GZ43_505278	U10400	Saccharomyces cerevisiae chromosome VIII cosmid L2825	1.80E-05
45	3544.A09.GZ43_505439	X75677	C.parapsilosis mt tRNA genes (591bps)	3.70E-08

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
46	3544.A13.GZ43_505503	D28811	Schistosoma japonicum mRNA for paramyosin, complete cds	5.40E-05
47	3544.A14.GZ43_505519	M87111	Human immunodeficiency virus type 2 (FORTC2) reverse transcriptase fragment	2.90E-05
48	3544.A17.GZ43_505567	L23650	Caenorhabditis elegans cosmid C27D11, complete sequence	5.60E-07
49	3544.B02.GZ43_505328	AF060543	Homo sapiens importin alpha 7 subunit mRNA, complete cds	1.50E-49
50	3544.B09.GZ43_505440	AB051473	Homo sapiens mRNA for KIAA1686 protein, partial cds	1.80E-05
51	3544.B18.GZ43_505584	AJ224821	Loxodonta africana complete mitochondrial genomic sequence	4.00E-06
52	3544.E05.GZ43_505379	AL451187	Human DNA sequence from clone RP11-49J23 on chromosome 6, complete sequence [Homo sapiens]	1.30E-07
53	3544.E18.GZ43_505587	L08338	Human immunodeficiency virus type 1 proviral envelope glycoprotein gene V3 region from A196/4537, clone 6 (from adult)	3.30E-07
54	3544.F06.GZ43_505396	X60833	R.norvegicus TDO2 gene for tryptophan 2,3-dioxygenase, exon 6	7.80E-07
55	3544.F16.GZ43_505556	U72716	Drosophila melanogaster D3-100EF mRNA, complete cds	2.00E-06
56	3544.G06.GZ43_505397	AC002359	Homo sapiens Xp22 Cosmid U239B3 (from Lawrence Livermore X library) complete sequence	1.60E-05
57	3544.G10.GZ43_505461	X56015	Crithidia oncopelti mitochondrial ND4, ND5, COI, 12S ribosomal RNA genes for NADH dehydrogenase subunit 4/5, cytochrome oxidase subunit I and 12S ribosomal RNA	4.80E-05
58	3544.G11.GZ43_505477	U80927	Dictyostelium discoideum unknown protein gene, complete cds	9.00E-08
59	3544.G12.GZ43_505493	AF245483	Oryza sativa OSE4 (OSE4) gene, complete cds	1.70E-07
60	3544.H03.GZ43_505350	Y12855	Homo sapiens P2X7 gene, exon 12 and 13	2.30E-05
61	3544.H15.GZ43_505542	AF194829	Tetragonia tetragonioides NADH dehydrogenase (ndhF) gene, partial cds; chloroplast gene for chloroplast product	2.00E-06
62	3544.H24.GZ43_505686	BC008353	Homo sapiens, Similar to RIKEN cDNA 0610008P16 gene, clone MGC:15937 IMAGE:3537224, mRNA, complete cds	2.50E-18
63	3544.I07.GZ43_505415	AF010533	Plasmodium falciparum microsatellite TA21 sequence	1.80E-08
64	3544.I15.GZ43_505543	D29794	Mouse gene for T cell receptor gamma chain	3.00E-06
65	3544.I20.GZ43_505623	AE000677	Aquifex aeolicus section 9 of 109 of the complete genome	4.00E-06
66	3544.J04.GZ43_505368	Z97015	Lactococcus lactis cremoris sucrose gene cluster	1.00E-06

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
67	3544.J11.GZ43_505480	M67480	Human prothymosin-alpha gene, complete cds	5.10E-10
68	3544.J13.GZ43_505512	AJ249884	Lepeophtheirus salmonis microsatellite DNA, locus Ls.NUIG.09	5.70E-08
69	3544.J23.GZ43_505672	AJ245823	Trypanosoma brucei PK4 gene for protein kinase	6.00E-06
70	3544.K16.GZ43_505561	U18191	Human HLA class I genomic survey sequence	2.50E-07
71	3544.L11.GZ43_505482	X07127	Kluyveromyces lactis killer plasmid k1 DNA	2.90E-05
72	3544.L13.GZ43_505514	BC005028	Homo sapiens, hypothetical protein FLJ11323, clone MGC:12582 IMAGE:3953383, mRNA, complete cds	1.80E-31
73	3544.M06.GZ43_505403	AC006687	Caenorhabditis elegans cosmid T20C7, complete sequence	2.30E-05
74	3544.M10.GZ43_505467	M92378	Mus musculus GABA transporter mRNA sequence	1.30E-05
75	3544.N07.GZ43_505420	U48705	Human receptor tyrosine kinase DDR gene, complete cds	7.40E-07
76	3544.N12.GZ43_505500	BC007621	Homo sapiens, Similar to Orthodenticle (Drosophila) homolog 1, clone MGC:15736 IMAGE:3355563, mRNA, complete cds	5.70E-07
77	3544.N19.GZ43_505612	AF270077	Staphylococcus epidermidis strain SR1 clone step.1047c06 genomic sequence	2.00E-07
78	3544.O03.GZ43_505357	U15681	Myrmecia pilosula H187-156 mitochondrion cytochrome b gene, partial cds	1.00E-06
79	3544.O10.GZ43_505469	AF056032	Homo sapiens kynurenine 3-hydroxylase mRNA, complete cds	5.00E-06
80	3544.O15.GZ43_505549	U37373	Xenopus laevis tail-specific thyroid hormone up-regulated (gene 5) mRNA, complete cds	3.00E-06
81	3544.O20.GZ43_505629	D66906	Bombyx mori DNA for sorbitol dehydrogenase, complete cds	2.00E-06
82	3544.P18.GZ43_505598	J04357	Red clover necrotic mosaic virus RNA-1, complete sequence	4.00E-06
83	3547.A04.GZ43_505743	AF118558	Mus musculus hitchhiker-3, hitchhiker-4, and hitchhiker-5 mRNA sequences	5.40E-07
84	3547.A11.GZ43_505855	U93874	Bacillus subtilis cysteine synthase (yrbA), cystathionine gamma-lyase (yrbB), YrbC (yrbC), YrbD (yrbD), formate dehydrogenase chain A (yrbE), YrbF (yrbF), formate dehydrogenase (yrbG), YrbH (yrbH), regulatory protein (yrbI), cytochrome P450 102 (yrbJ), >	4.00E-06
85	3547.A24.GZ43_506063	AL157466	Homo sapiens mRNA; cDNA DKFZp761E2423 (from clone DKFZp761E2423)	8.80E-07
86	3547.C05.GZ43_505761	X52589	Bovine rotavirus RNA for virus protein 2 (VP2)	1.00E-05



Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
87	3547.C17.GZ43_505953	U67594	<i>Methanococcus jannaschii</i> section 136 of 150 of the complete genome	3.80E-05
88	3547.C23.GZ43_506049	AJ250862	<i>Bacillus</i> sp. HIL-Y85/54728 mersacidin biosynthesis gene cluster (mrsK2, mrsR2, mrsF, mrsG, mrsE, mrsA, mrsR1, mrsD, mrsM and mrsT genes)	1.20E-05
89	3547.D19.GZ43_505986	AF050491	<i>Microgadus tomcod</i> aromatic hydrocarbon receptor (ahr) gene, exons 8-11, partial cds	4.00E-06
90	3547.D23.GZ43_506050	M33190	Rat cytochrome P450 II A3 (CYP2A3) gene, complete cds	5.80E-05
91	3547.E04.GZ43_505747	L35658	Homo sapiens (subclone H8 9_d12 from P1 35 H5 C8) DNA sequence	7.70E-07
92	3547.F02.GZ43_505716	AF038190	Homo sapiens clone 23582 mRNA sequence	1.10E-07
93	3547.F10.GZ43_505844	AY008833	<i>Staphylococcus aureus</i> tcaR-tcaA-tcaB operon, complete sequences	5.00E-06
94	3547.F20.GZ43_506004	AB037821	Homo sapiens mRNA for KIAA1400 protein, partial cds	1.00E-06
95	3547.G02.GZ43_505717	M88397	<i>Naegleria fowleri</i> virulence-related protein (NF314) mRNA, complete cds	3.70E-07
96	3547.G09.GZ43_505829	AJ315644	Homo sapiens mRNA for proton myo-inositol symporter (Hmit gene)	7.90E-07
97	3547.G22.GZ43_506037	Z33603	<i>P.radiata</i> (Pr1.6) microsatellite DNA, 703bp	1.70E-07
98	3547.H12.GZ43_505878	L04309	<i>Shigella flexneri</i> ipgD, ipgE, ipgF genes, complete cds	3.00E-06
99	3547.H14.GZ43_505910	AL137502	Homo sapiens mRNA; cDNA DKFZp761H171 (from clone DKFZp761H171); partial cds	2.90E-07
100	3547.I07.GZ43_505799	M15332	<i>B.sphaericus</i> ermG gene encoding rRNA methyltransferase (macrolide-lincosamide-streptogramin B resistance element)	7.00E-06
101	3547.I16.GZ43_505943	AF015157	Homo sapiens clone HS19.12 Alu-Ya5 sequence	4.70E-10
102	3547.I17.GZ43_505959	AE007758	<i>Clostridium acetobutylicum</i> ATCC824 section 246 of 356 of the complete genome	3.00E-06
103	3547.I20.GZ43_506007	L37606	<i>Medicago sativa</i> (clone GG16-1) NADH-dependent glutamate synthase gene, complete cds	1.50E-05
104	3547.J05.GZ43_505768	Z16911	H. sapiens (D20S113) DNA segment containing (CA) repeat; clone AFM205th8, single read	2.80E-07
105	3547.J10.GZ43_505848	Z37803	HIV-1 DNA V3 region (patient 15, sample CSF, clone 9)	8.80E-07
106	3547.J20.GZ43_506008	AF013273	<i>Candida albicans</i> histidine kinase 1 gene, complete cds	3.30E-05
107	3547.J22.GZ43_506040	AF289080	<i>Lycopersicon esculentum</i> alpha-galactosidase gene, partial cds	4.00E-06

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
108	3547.K01.GZ43_505705	AF267863	Homo sapiens DC43 mRNA, complete cds	7.30E-22
109	3547.L09.GZ43_505834	Z22175	Caenorhabditis elegans cosmid K01F9, complete sequence	1.40E-05
110	3547.L11.GZ43_505866	AJ288648	Limnodynastes tasmaniensis mitochondrial partial nadh4 gene for NADH dehydrogenase subunit 4 and partial tRNA-His gene, sample 26 from Australia Boolara	5.90E-07
111	3547.L16.GZ43_505946	AE001293	Chlamydia trachomatis section 20 of 87 of the complete genome	7.10E-07
112	3547.L22.GZ43_506042	AF287006	Danio rerio T-box brain 1 mRNA, partial cds	7.00E-06
113	3547.M02.GZ43_505723	AE007788	Clostridium acetobutylicum ATCC824 section 276 of 356 of the complete genome	1.00E-05
114	3547.M07.GZ43_505803	Z46252	M.musculus DNA for region surrounding retrovirus restriction locus Fv1	6.00E-06
115	3547.M08.GZ43_505819	AB020684	Homo sapiens mRNA for KIAA0877 protein, partial cds	1.50E-05
116	3547.M16.GZ43_505947	AF335240	Petunia x hybrida MADS-box transcription factor FBP22 (FBP22) mRNA, complete cds	3.00E-06
117	3547.N06.GZ43_505788	AF299346	Remispora flavissima isolate CEH313 18S ribosomal RNA gene, partial sequence; internal transcribed spacer 1, 5.8S ribosomal RNA gene and internal transcribed spacer 2, complete sequence; and 28S ribosomal RNA gene, partial sequence	1.70E-08
118	3547.O03.GZ43_505741	AE002344	Chlamydia muridarum, section 72 of 85 of the complete genome	6.60E-07
119	3547.O07.GZ43_505805	D50608	Rat gene for cholecystokinin type-A receptor (CCKAR), complete cds	1.60E-05
120	3547.O14.GZ43_505917	AL137502	Homo sapiens mRNA; cDNA DKFZp761H171 (from clone DKFZp761H171); partial cds	2.90E-07
121	3547.P18.GZ43_505982	AJ131734	Plasmodium berghei DNA including upstream sequence NTS and 5'ETS of the 18S rRNA gene (A rRNA gene unit)	6.10E-07
122	3547.P21.GZ43_506030	AC006619	Caenorhabditis elegans cosmid C46C11, complete sequence	1.70E-05
123	3547.P22.GZ43_506046	AJ000871	Streptococcus mitis comC, comD, comE genes, isolate B5	2.00E-06
124	3550.A12.GZ43_506255	M22310	Human epidermal growth factor receptor proto-oncogene downstream enhancer	4.80E-07
125	3550.A16.GZ43_506319	L39435	Senecio mikanioides chloroplast NADH dehydrogenase (ndhF) gene, complete cds	2.00E-06
126	3550.B06.GZ43_506160	D14161	Hordeum vulgare ids-4 mRNA, complete cds	1.10E-08

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
127	3550.C01.GZ43_506081	AK025182	Homo sapiens cDNA: FLJ21529 fis, clone COL05981	4.20E-09
128	3550.C22.GZ43_506417	X52028	Rattus norvegicus P450 IID3 gene	1.41E-04
129	3550.D16.GZ43_506322	Y10345	H.sapiens GalNAc-T3 gene, 3'UTR	5.00E-07
130	3550.D23.GZ43_506434	AF134403	Escherichia coli plasmid pAA2 Shf (shf), hexosyltransferase homolog (capU), and VirK (virK) genes, complete cds	6.90E-07
131	3550.E02.GZ43_506099	U66074	Trilichomonas foetus putative superoxide dismutase 2 (SOD2) gene, complete cds	8.90E-07
132	3550.E06.GZ43_506163	Z23341	H. sapiens (D8S528) DNA segment containing (CA) repeat; clone AFM080sh7; single read	2.30E-08
133	3550.F06.GZ43_506164	M59447	Drosophila melanogaster Sex-lethal (Sxl) mRNA, complete cds	3.00E-06
134	3550.F08.GZ43_506196	M24901	Rabbit pulmonary surfactant-associated protein (SP-B) mRNA, complete cds	3.40E-07
135	3550.F20.GZ43_506388	AF216169	Simicratea welwitschii clone 2 phytochrome B (PHYB) gene, exon 1 and partial cds	5.40E-08
136	3550.F22.GZ43_506420	AP000739	Arabidopsis thaliana genomic DNA, chromosome 3, P1 clone:MEK6	2.20E-05
137	3550.G02.GZ43_506101	AL022342	Human DNA sequence from clone RP1-29M10 on chromosome 20, complete sequence [Homo sapiens]	7.40E-05
138	3550.G08.GZ43_506197	AK021312	Mus musculus 13 days embryo stomach cDNA, RIKEN full-length enriched library, clone:D530039A21, full insert sequence	3.60E-08
139	3550.G10.GZ43_506229	M31684	D.melanogaster cytoskeleton-like bicaudalD protein (BicD) mRNA, complete cds	3.00E-06
140	3550.G15.GZ43_506309	AF087141	Mus musculus uncharacterized long terminal repeat, complete sequence; and valyl-tRNA synthetase (G7a) gene, complete cds	4.00E-06
141	3550.G23.GZ43_506437	X02547	Trypanosoma brucei mitochondrial genes for 12S and 9S ribosomal RNA	2.00E-06
142	3550.H10.GZ43_506230	U55711	Human ataxia-telangiectasia (ATM) gene, exon 11	6.10E-08
143	3550.H21.GZ43_506406	Z68755	Human DNA sequence from cosmid L118D5, Huntington's Disease Region, chromosome 4p16.3	2.00E-06
144	3550.H23.GZ43_506438	AF151388	Dermatobia hominis strain Alienus tRNA-Ile gene, partial sequence; D-loop, complete sequence; and 12S ribosomal RNA, partial sequence; mitochondrial genes for mitochondrial products	1.20E-07

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
145	3550.I03.GZ43_506119	AF117258	Staphylococcus aureus plasmid pIP680 replication protein RepE (repE) gene, partial cds; resolvase (res), acetyltransferase Vat (vat), and hydrolase VgB (vgb) genes, complete cds; and unknown gene	6.50E-08
146	3550.I19.GZ43_506375	AE002781	Drosophila melanogaster genomic scaffold 142000013385442, complete sequence	3.90E-05
147	3550.I21.GZ43_506407	AE001002	Archaeoglobus fulgidus section 105 of 172 of the complete genome	4.20E-05
148	3550.J05.GZ43_506152	AF080689	Homo sapiens protein kinase PITSLRE (CDC2L2) gene, exons 8 and 9	5.50E-10
149	3550.J11.GZ43_506248	Z82761	R.prowazekii genomic DNA fragment (clone A793R)	1.00E-06
150	3550.K05.GZ43_506153	X15407	Maize pseudo-Gpa2 pseudogene for glyceraldehyde-3-phosphate dehydrogenase subunit A	3.20E-05
151	3550.K09.GZ43_506217	X62631	S.pombe wis1 gene for protein kinase	1.50E-07
152	3550.K14.GZ43_506297	M59743	Rabbit cardiac muscle Ca <sup>2+</sup> release channel (ryanodine receptor) mRNA, complete cds	1.00E-06
153	3550.L16.GZ43_506330	AF201383	Buchnera aphidicola isopropylmalate dehydratase subunit (IeuC) gene, partial cds	1.00E-06
154	3550.L19.GZ43_506378	M77244	H.sapiens erythropoietin receptor (EPOR) gene, 5' end	4.00E-09
155	3550.L23.GZ43_506442	L76259	Homo sapiens PTS gene, complete cds	8.00E-06
156	3550.M21.GZ43_506411	M87339	Human replication factor C, 37-kDa subunit mRNA, complete cds	5.00E-06
157	3550.N01.GZ43_506092	AF191009	Helicobacter pylori strain ChinaF30A cag pathogenicity island polymorphic right end, type IIIa motif	1.10E-07
158	3550.N07.GZ43_506188	AF235499	Mus musculus SH2-containing inositol 5-phosphatase (Ship) gene, exons 3 through 6	1.55E-04
159	3550.O03.GZ43_506125	D14813	Human DNA for osteopontin, complete cds	4.50E-05
160	3550.O04.GZ43_506141	U08596	Canis familiaris delayed rectifier K <sup>+</sup> channel mRNA, partial cds	6.00E-06
161	3550.O08.GZ43_506205	XM_017044	Homo sapiens similar to diaphanous (Drosophila, homolog) 2 (H. sapiens) (LOC91459), mRNA	6.40E-09
162	3550.O15.GZ43_506317	U15977	Mus musculus long chain fatty acyl CoA synthetase mRNA, complete cds	2.80E-05
163	3550.O17.GZ43_506349	X62578	C. caldarium plastid genes ompR', psbD, psbC, rps16 and groEL	2.80E-05
164	3550.O18.GZ43_506365	L34363	Human X-linked nuclear protein (XNP) gene, complete cds	4.00E-06
165	3550.O21.GZ43_506413	AB056784	Macaca fascicularis brain cDNA clone:QnpA-11501, full insert sequence	5.20E-07

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
166	3550.P18.GZ43_506366	AK002041	Homo sapiens cDNA FLJ11179 fis, clone PLACE1007450	5.30E-07
167	3550.P23.GZ43_506446	AF200361	Rattus norvegicus cytochrome P450 4F1 (Cyp4F1) gene, complete cds	1.40E-05
168	3553.A09.GZ43_506591	AL109980	Human DNA sequence from clone RP4-697G8 on chromosome 22, complete sequence [Homo sapiens]	3.50E-12
169	3553.B07.GZ43_506560	L37056	Strongylocentrotus purpuratus myc protein mRNA, complete cds	4.60E-07
170	3553.B16.GZ43_506704	U43542	Nicotiana tabacum diphenol oxidase mRNA, complete cds	2.00E-06
171	3553.B22.GZ43_506800	L34040	Homo sapiens stromelysin gene, promoter region	6.00E-06
172	3553.D04.GZ43_506514	Y07599	S.pombe mRNA for dmf1 gene	9.40E-07
173	3553.D07.GZ43_506562	X13835	R.norvegicus CaMII gene, exons 3,4 & 5	2.00E-06
174	3553.D14.GZ43_506674	L38424	Bacillus subtilis dihydronicotinamide reductase (jojE) gene, complete cds; poly(A) polymerase (jojI) gene, complete cds; biotin acetyl-CoA-carboxylase ligase (birA) gene, complete cds; jojC, jojD, jojF, jojG, jojH genes, complete cds's	1.80E-05
175	3553.D19.GZ43_506754	X53431	Yeast gene for STE11	9.00E-06
176	3553.E08.GZ43_506579	AF062863	Arabidopsis thaliana putative transcription factor (MYB11) mRNA, partial cds	1.80E-07
177	3553.E09.GZ43_506595	X71067	X.laevis XFG 5-1 and XFG 5-2 genes for zinc finger proteins	6.60E-05
178	3553.F12.GZ43_506644	X63223	B.taurus Cl-MNLL mRNA for ubiquinone oxidoreductase complex	6.90E-08
179	3553.F13.GZ43_506660	L81869	Homo sapiens (subclone 1_c4 from P1 H55) DNA sequence, complete sequence	3.00E-08
180	3553.F19.GZ43_506756	U97190	Caenorhabditis elegans cosmid B0025, complete sequence	3.00E-06
181	3553.G05.GZ43_506533	S76404	beta -HKA-H,K-ATPase beta-subunit [rats, Genomic, 8983 nt, segment 2 of 2]	8.00E-06
182	3553.G06.GZ43_506549	X68048	Phaseolus vulgaris chloroplast DNA for tRNA-His gene region	5.00E-06
183	3553.G07.GZ43_506565	AF068289	Homo sapiens HDCMD34P mRNA, complete cds	4.40E-12
184	3553.G21.GZ43_506789	Z33603	P.radiata (Pr1.6) microsatellite DNA, 703bp	1.70E-07
185	3553.H06.GZ43_506550	AF090901	Homo sapiens clone HQ0195\$ PRO0195 mRNA, complete cds	8.00E-07
186	3553.H09.GZ43_506598	AF270105	Staphylococcus epidermidis strain SR1 clone step.1049c09 genomic sequence	9.80E-07
187	3553.H21.GZ43_506790	Z18359	Glycine max seed-specific low molecular weight sulfur-rich protein	2.00E-06
188	3553.I13.GZ43_506663	AF155115	Homo sapiens NY-REN-58 antigen mRNA, complete cds	1.70E-07

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
189	3553.I16.GZ43_506711	AF270229	Staphylococcus epidermidis strain SR1 clone step.1055d10 genomic sequence	1.20E-05
190	3553.J12.GZ43_506648	U53400	Rattus norvegicus chromosome 10 microsatellite sequence D10Mco21	1.55E-01
191	3553.J14.GZ43_506680	M10014	Homo sapiens map 4q28 fibrinogen (FGB) gene, alternative splice products, complete cds	9.00E-06
192	3553.J16.GZ43_506712	Z23341	H. sapiens (D8S528) DNA segment containing (CA) repeat; clone AFM080xh7; single read	2.30E-08
193	3553.J17.GZ43_506728	AK021312	Mus musculus 13 days embryo stomach cDNA, RIKEN full-length enriched library, clone:D530039A21, full insert sequence	3.60E-08
194	3553.J22.GZ43_506808	AF120279	Mus musculus proline dehydrogenase mRNA, complete cds	5.00E-06
195	3553.J24.GZ43_506840	Z18359	Glycine max seed-specific low molecular weight sulfur-rich protein	2.00E-06
196	3553.K01.GZ43_506473	U31465	Kluyveromyces lactis telomerase RNA component (TER1) gene, complete sequence	2.00E-06
197	3553.K02.GZ43_506489	X60672	M.musculus mRNA for radixin	1.00E-06
198	3553.K03.GZ43_506505	Z71943	G.hyalina (92-89) DNA for internal transcribed spacer 1	1.06E-02
199	3553.K05.GZ43_506537	M80596	Saccharomyces cerevisiae VAC1 gene (required for vacuole inheritance and vacuole protein sorting), complete cds	6.00E-06
200	3553.K07.GZ43_506569	AJ275317	Cicer arietinum partial mRNA for malate dehydrogenase	7.60E-07
201	3553.K15.GZ43_506697	X57377	Mouse dilute myosin heavy chain gene for novel heavy chain with unique C-terminal region	2.40E-05
202	3538.A11.GZ43_504703	Z75199	S.cerevisiae chromosome XV reading frame ORF YOR291w	6.00E-06
203	3538.A24.GZ43_504911	AF270077	Staphylococcus epidermidis strain SR1 clone step.1047c06 genomic sequence	2.00E-07
204	3538.B01.GZ43_504544	AF368255	Arabidopsis thaliana small zinc finger-like protein TIM13 mRNA, complete cds; nuclear gene for mitochondrial product	4.10E-07
205	3538.B20.GZ43_504848	AB069994	Macaca fascicularis testis cDNA clone:QtsA.10636, full insert sequence	1.40E-07
206	3538.C01.GZ43_504545	AF072375	Pseudoalteromonas sp. S9 beta-hexosaminidase (chiP) gene, complete cds	1.50E-04
207	3538.C02.GZ43_504561	AJ011271	Human immunodeficiency virus type 2 partial env gene, isolate b1286	7.10E-08
208	3538.D06.GZ43_504626	AF100765	Oryza sativa receptor-like kinase (SARK1) gene, complete cds	3.00E-06
209	3538.D09.GZ43_504674	Z47784	M.musculus mRNA expressed in islet cells (clone 58)	3.40E-08

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
210	3538.D21.GZ43_504866	AE006568	<i>Streptococcus pyogenes</i> M1 GAS strain SF370, section 97 of 167 of the complete genome	6.00E-07
211	3538.E15.GZ43_504771	AB027966	<i>Schizosaccharomyces pombe</i> gene for Hypothetical protein, partial cds, clone:TB89	2.30E-08
212	3538.F02.GZ43_504564	AK001871	Homo sapiens cDNA FLJ11009 fis, clone PLACE1003108	5.90E-09
213	3538.F08.GZ43_504660	U39161	Human phosphodiesterase (PDEA) gene, intron 8, 5' end	7.40E-07
214	3553.K23.GZ43_506825	Y12052	Homo sapiens gene encoding guanine nucleotide-binding protein beta3 subunit, exon 5	3.00E-06
215	3553.K24.GZ43_506841	AP001419	Homo sapiens genomic DNA, chromosome 21q22.2, clone:PAC24K9, LB7T-ERG region, complete sequence	1.00E-06
216	3553.L02.GZ43_506490	X15028	Chicken hsp90 gene for 90 kDa-heat shock protein 5'-end	3.60E-05
217	3553.L04.GZ43_506522	L34665	<i>Rattus norvegicus</i> H+/K+-ATPase beta subunit (HKB) gene, exon 6	1.20E-09
218	3553.L21.GZ43_506794	M87843	Human transforming growth factor beta-2 gene, 5' end	2.30E-05
219	3553.M12.GZ43_506651	AB026592	<i>Limnoporus caekii</i> mitochondrial gene for 16S ribosomal RNA, partial sequence	1.10E-07
220	3553.M23.GZ43_506827	AE006349	<i>Lactococcus lactis</i> subsp. <i>lactis</i> IL1403 section 111 of 218 of the complete genome	8.00E-07
221	3553.N01.GZ43_506476	U80457	Human transcription factor SIM2 short form mRNA, complete cds	2.00E-06
222	3553.N02.GZ43_506492	U81144	<i>Caenorhabditis elegans</i> non-alpha nicotinic acetylcholine receptor subunit precursor (unc-29) gene, complete cds	3.20E-07
223	3553.N04.GZ43_506524	AE006296	<i>Lactococcus lactis</i> subsp. <i>lactis</i> IL1403 section 58 of 218 of the complete genome	2.00E-06
224	3553.N07.GZ43_506572	AK026258	Homo sapiens cDNA: FLJ22605 fis, clone HSI04743	1.00E-06
225	3553.N08.GZ43_506588	U05822	Human proto-oncogene BCL3 gene, exon 2	1.90E-14
226	3553.O07.GZ43_506573	X97196	<i>D.melanogaster</i> X gene	4.00E-06
227	3553.O18.GZ43_506749	AE001146	<i>Borrelia burgdorferi</i> (section 32 of 70) of the complete genome	1.60E-05
228	3553.O23.GZ43_506829	X63509	<i>Mus musculus</i> partial L1 gene, exons 2-4	6.00E-06
229	3553.P03.GZ43_506510	M24901	Rabbit pulmonary surfactant-associated protein (SP-B) mRNA, complete cds	4.20E-07
230	3553.P05.GZ43_506542	AF239156	Homo sapiens peptide deformylase-like protein mRNA, complete cds	1.00E-06
231	3553.P12.GZ43_506654	AF283753	<i>Acipenser persicus</i> isolate cw203 cytochrome b gene, partial cds; mitochondrial gene for mitochondrial product	3.90E-07

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
232	3553.P18.GZ43_506750	AK021312	Mus musculus 13 days embryo stomach cDNA, RIKEN full-length enriched library, clone:D530039A21, full insert sequence	3.50E-08
233	3553.P21.GZ43_506798	AB044136	Homo sapiens genomic DNA, clone:#7	4.00E-06
234	3556.A03.GZ43_506879	X61084	C.griscus rhodopsin gene for opsin protein	4.30E-05
235	3556.A06.GZ43_506927	L46904	Homo sapiens (subclone 4_c6 from P1 H22) DNA sequence	1.20E-08
236	3556.B06.GZ43_506928	AK002041	Homo sapiens cDNA FLJ11179 fis, clone PLACE1007450	1.40E-07
237	3556.B09.GZ43_506976	U88832	Human groucho protein homolog (AES) gene, exons 2-7 and complete cds	7.00E-07
238	3556.B10.GZ43_506992	M11925	Influenza A/chicken/Pennsylvania/8125/83 (H5N2) neuraminidase (NA) gene, complete cds	5.00E-06
239	3556.B14.GZ43_507056	Z80218	Caenorhabditis elegans cosmid F52D4, complete sequence	2.20E-05
240	3556.C13.GZ43_507041	AF348512	Mus musculus polyamine-modulated factor-1 gene, exons 2 through 5 and complete cds	8.00E-06
241	3556.C15.GZ43_507073	X82013	S.cerevisiae mRNA for SUL1	3.00E-06
242	3556.C18.GZ43_507121	Z23973	H. sapiens (D7S660) DNA segment containing (CA) repeat; clone AFM277vd5; single read	5.00E-06
243	3556.C24.GZ43_507217	AE001381	Plasmodium falciparum chromosome 2, section 18 of 73 of the complete sequence	6.90E-07
244	3556.D15.GZ43_507074	U48288	Rattus norvegicus A-kinase anchoring protein AKAP 220 mRNA, complete cds	5.50E-07
245	3556.D20.GZ43_507154	AF092684	Neochlamis scabripennis haplotype 113 cytochrome oxidase I (COI) gene, mitochondrial gene encoding mitochondrial protein, partial cds	4.00E-07
246	3556.D23.GZ43_507202	X16416	Human c-abl mRNA encoding p150 protein	2.25E-04
247	3556.E13.GZ43_507043	AL049948	Homo sapiens mRNA; cDNA DKFPz564K0222 (from clone DKFPz564K0222)	6.60E-08
248	3556.E24.GZ43_507219	Z57634	H.sapiens CpG island DNA genomic MseI fragment, clone 187e9, forward read cpg187e9.f11a	8.70E-07
249	3556.F10.GZ43_506996	AF025409	Homo sapiens zinc transporter 4 (ZNT4) mRNA, complete cds	3.90E-34
250	3556.G15.GZ43_507077	X15407	Maize pseudo-Gpa2 pseudogene for glyceraldehyde-3-phosphate dehydrogenase subunit A	3.40E-05
251	3556.H01.GZ43_506854	AF269443	Staphylococcus epidermidis strain SR1 clone step.1003h04 genomic sequence	3.00E-06



Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
252	3556.H02.GZ43_506870	U31465	Kluyveromyces lactis telomerase RNA component (TER1) gene, complete sequence	3.00E-06
253	3556.H12.GZ43_507030	Z68886	Human DNA sequence from cosmid L21F12, Huntington's Disease Region, chromosome 4p16.3	1.70E-07
254	3556.H20.GZ43_507158	AB034628	Equus caballus microsatellite TKY319, TKY320 DNA	1.70E-07
255	3556.I02.GZ43_506871	AL390767	Human DNA sequence from clone RP1-68P15 on chromosome 11p13-14.2 Contains GSSs and ESTs. Contains part of a novel gene, complete sequence [Homo sapiens]	2.00E-06
256	3556.I14.GZ43_507063	U34042	Mus musculus mammalian tolloid-like protein mRNA, complete cds	1.50E-05
257	3556.J05.GZ43_506920	U31465	Kluyveromyces lactis telomerase RNA component (TER1) gene, complete sequence	2.00E-06
258	3556.J07.GZ43_506952	AL359621	Homo sapiens mRNA; cDNA DKFZp434M1631 (from clone DKFZp434M1631)	2.00E-06
259	3556.J14.GZ43_507064	M81830	Human somatostatin receptor isoform 2 (SSTR2) gene, complete cds	1.00E-06
260	3556.J16.GZ43_507096	Y15484	Canis familiaris gene encoding retinal guanylate cyclase E	2.90E-08
261	3556.K04.GZ43_506905	U88832	Human groucho protein homolog (AES) gene, exons 2-7 and complete cds	8.00E-07
262	3556.K12.GZ43_507033	AP001419	Homo sapiens genomic DNA, chromosome 21q22.2, clone:PAC24K9, LB7T-ERGG region, complete sequence	1.00E-06
263	3556.K13.GZ43_507049	AK023589	Homo sapiens cDNA FLJ13527 fis, clone PLACE1006076	2.00E-06
264	3556.K17.GZ43_507113	X71634	D.bifasciata P-Transposon	3.00E-06
265	3556.L08.GZ43_506970	X02367	Glaucoma chattoni rDNA 3' NTS	8.20E-08
266	3556.L09.GZ43_506986	AF154329	Pisum sativum MAP kinase PsMAPK2 (Mapk2) mRNA, complete cds	4.10E-07
267	3556.L16.GZ43_507098	AB041791	Homo sapiens HSPDE10A gene for phosphodiesterase 10A1 (PDE10A1), exon 17	3.10E-08
268	3556.L23.GZ43_507210	M23720	Rat carboxypeptidase (CA2) gene, exon 10	5.00E-06
269	3556.M02.GZ43_506875	U91963	Human tolloid-like protein (TLL) mRNA, complete cds	1.40E-05
270	3556.M11.GZ43_507019	X16353	Rickettsia ompB gene for outer membrane protein B	7.60E-05
271	3556.M23.GZ43_507211	X93496	H.sapiens TRAP gene, 5' flanking region	5.60E-23
272	3556.N02.GZ43_506876	U26458	Snakehead retrovirus (SnRV), complete genome	3.20E-05
273	3556.N04.GZ43_506908	L39064	Homo sapiens interleukin 9 receptor precursor (IL9R) gene, complete cds	4.00E-09

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
274	3556.N05.GZ43_506924	M63437	Chicken KLG gene, complete cds	2.00E-06
275	3556.N06.GZ43_506940	AF327424	Arabidopsis thaliana unknown protein (T14P1.19/At2g45010) mRNA, partial cds	2.00E-07
276	3556.N21.GZ43_507180	AB022157	Mus musculus Cctd gene for chaparonin containing TCP-1 delta subunit, complete cds	4.00E-06
277	3556.O08.GZ43_506973	X00171	Vibrio cholera toxin (ctx) operon DNA sequence from strain 2125	7.00E-06
278	3556.O13.GZ43_507053	U41106	Caenorhabditis elegans cosmid W06A11	1.20E-05
279	3556.P07.GZ43_506958	M15085	T.brucei expressed copy of the ILTat 1.3 variable surface glycoprotein gene, 5' flank	1.10E-07
280	3559.A04.GZ43_507279	AE006824	Sulfolobus solfataricus section 183 of 272 of the complete genome	4.70E-05
281	3559.A20.GZ43_507535	X71787	A.thaliana AAP2 mRNA for amino acid permease	2.00E-06
282	3559.A24.GZ43_507599	X56494	H.sapiens M gene for M1-type and M2-type pyruvate kinase	1.80E-05
283	3559.B04.GZ43_507280	AJ215550	Homo sapiens partial AK155 gene for AK155 protein, exons 1-3 and joined CDS	2.50E-05
284	3559.B06.GZ43_507312	AF077344	Homo sapiens cartilage-derived C-type lectin (CLECSF1) gene, exons 1 and 2	5.80E-05
285	3559.B08.GZ43_507344	D50552	Xenopus laevis xSox12 mRNA for XSOX12, complete cds	4.00E-07
286	3559.B10.GZ43_507376	L76259	Homo sapiens PTS gene, complete cds	9.00E-06
287	3559.B18.GZ43_507504	M29109	D.discoideum actin M6 gene, 5' flank	3.40E-07
288	3559.C06.GZ43_507313	X99910	C.carpio mRNA transcription factor, ovxl	1.60E-05
289	3559.D21.GZ43_507554	AK022877	Homo sapiens cDNA FLJ12815 fls, clone NT2RP2002546	2.00E-06
290	3559.E06.GZ43_507315	U97408	Caenorhabditis elegans cosmid F48A9	3.00E-06
291	3559.E09.GZ43_507363	L40489	Ureaplasma urealyticum UreA (ureA), UreB (ureB), UreC (ureC), UreE (ureE), UreF (ureF), and UreG (ureG) genes, complete cds; UreD (ureD) gene, partial cds; and unknown gene	3.00E-07
292	3559.E20.GZ43_507539	AF113521	Zea mays putative transcription factor mRNA sequence	8.20E-08
293	3559.F07.GZ43_507332	AF109377	Mus musculus Idl1p (LDLB) mRNA, complete cds	4.30E-05
294	3559.F17.GZ43_507492	U11292	Human Ki nuclear autoantigen mRNA, complete cds	6.40E-07
295	3559.H09.GZ43_507366	X13414	Murine I gene for MHC class II(Ia) associated invariant chain	9.00E-06
296	3559.H22.GZ43_507574	U61402	Streptococcus thermophilus GalR (galR), galactokinase (galK) and gal-1-P uridylyltransferase (galT) genes, complete cds	1.00E-06
297	3559.H24.GZ43_507606	U67594	Methanococcus jannaschii section 136 of 150 of the complete genome	3.40E-05
298	3559.I05.GZ43_507303	X97289	S.salar genes encoding alpha-globin and beta-globin, clone 6	7.00E-06

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
299	3559.J04.GZ43_507288	L10709	Human constitutive endothelial nitric oxide synthase gene, exons 25 and 26 and complete cds	8.90E-12
300	3559.J20.GZ43_507544	U67559	Methanococcus jannaschii section 101 of 150 of the complete genome	5.70E-05
301	3559.K16.GZ43_507481	Z48955	D.virginiana partial LINE-1 repetitive DNA and putative RT	2.40E-08
302	3559.K17.GZ43_507497	AC004497	Homo sapiens chromosome 21, P1 clone LBNL#6 (LBNL H10), complete sequence	4.00E-06
303	3559.L01.GZ43_507242	X58774	Herpesvirus saimiri sRNA1, sRNA2, sRNA3 and sRNA4 genes for small viral RNAs	1.00E-06
304	3559.L14.GZ43_507450	X67774	C.upsaliensis (LMG 8854) 23S rRNA gene	1.30E-05
305	3559.L19.GZ43_507530	Z57634	H.sapiens CpG island DNA genomic MseI fragment, clone 187e9, forward read cpgl87e9.fl1a	7.70E-07
306	3559.M02.GZ43_507259	AF042834	Homo sapiens phosphodiesterase delta subunit gene, exons 2, 3 and 4	1.30E-05
307	3559.M09.GZ43_507371	U07628	Caenorhabditis elegans N2 APX-1 (apx-1) mRNA, complete cds	2.00E-06
308	3559.N05.GZ43_507308	Z24259	H. sapiens (D19S417) DNA segment containing (CA) repeat; clone AFM304zg1; single read	3.70E-07
309	3559.N18.GZ43_507516	S75829	{dinucleotide repeats, microsatellite marker} [Dryobalanops lanceolata, Genomic, 230 nt]	1.90E-07
310	3559.N21.GZ43_507564	AL353948	Homo sapiens mRNA; cDNA DKFZp761P0114 (from clone DKFZp761P0114)	5.30E-07
311	3559.O01.GZ43_507245	AL110269	Homo sapiens mRNA; cDNA DKFZp564A122 (from clone DKFZp564A122); partial cds	1.60E-17
312	3559.O05.GZ43_507309	Y08695	Clostridium tertium nanH gene	7.40E-07
313	3559.O07.GZ43_507341	AJ249489	Xenopus laevis partial mRNA for putative olfactory receptor (xb6 gene)	5.40E-07
314	3559.O20.GZ43_507549	X02886	Human gene for T-cell receptor alpha chain J region	2.00E-06
315	3559.P10.GZ43_507390	X66030	Homo sapiens partial ufo gene encoding tyrosine kinase receptor	4.90E-07
316	3559.P15.GZ43_507470	Z16777	H. sapiens (D2S139) DNA segment containing (CA) repeat; clone AFM177xh4; single read	4.00E-06
317	3559.P18.GZ43_507518	AJ228072	Nicotiana benthamiana DNA for Tnt1 retrotransposable element, isolate ben15	2.80E-07
318	3559.P24.GZ43_507614	U32372	Rattus norvegicus tyrosine-ester sulfotransferase mRNA, complete cds	4.90E-07
319	3562.A01.GZ43_507615	AE000496	Escherichia coli K12 MG1655 section 386 of 400 of the complete genome	1.56E-04
320	3562.A15.GZ43_507839	AF068289	Homo sapiens HDCMD34P mRNA, complete cds	6.60E-11

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
321	3562.B22.GZ43_507952	AK014534	Mus musculus 0 day neonate skin cDNA, RIKEN full-length enriched library, clone:4631424J17, full insert sequence	1.10E-07
322	3562.C23.GZ43_507969	L01787	Ascaris sumum phosphoenolpyruvate carboxykinase (PEPCK) gene, complete cds	3.70E-07
323	3562.D10.GZ43_507762	X54061	D. melanogaster mRNA coding for a 205K microtubule-associated protein (MAP)	6.60E-07
324	3562.E01.GZ43_507619	M29812	Homo sapiens Ig H-chain V71-4 (IGH@) gene, partial cds	1.50E-05
325	3562.E03.GZ43_507651	X03729	Vaccinia virus late gene cluster from central portion of genome containing the L65 gene locus	2.63E-04
326	3562.E12.GZ43_507795	M29694	B.licheniformis RNA polymerase sigma-30 factor (spo0H) gene, complete cds	1.60E-05
327	3562.F19.GZ43_507908	AE006216	Pasteurella multocida PM70 section 183 of 204 of the complete genome	2.40E-05
328	3562.F20.GZ43_507924	M91004	Rabbit endothelial leukocyte adhesion molecule I (ELAMI), complete cds	2.00E-06
329	3562.G13.GZ43_507813	X69818	E.muelleri COLF1 gene for extracellular matrix protein	1.00E-06
330	3562.G19.GZ43_507909	AK019034	Mus musculus 10 day old male pancreas cDNA, RIKEN full-length enriched library, clone:1810049K24, full insert sequence	1.00E-05
331	3562.H11.GZ43_507782	AF206598	Algyroides fitzingeri 12S ribosomal RNA gene, partial sequence; tRNA-Val gene, complete sequence; and 16S ribosomal RNA gene, partial sequence; mitochondrial genes for mitochondrial products	1.40E-07
332	3562.H12.GZ43_507798	AK002041	Homo sapiens cDNA FLJ11179 fis, clone PLACE1007450	5.30E-07
333	3562.I01.GZ43_507623	S39048	knob associated histidine-rich protein KAHRP (5'region) [Plasmodium falciparum, Genomic, 2215 nt]	2.00E-06
334	3562.I02.GZ43_507639	AF129501	Buchnera aphidicola natural-host Diuraphis noxia acetohydroxy acid synthase large subunit (ilvI) and acetohydroxy acid synthase small subunit (ilvH) genes, complete cds; and unknown genes	1.60E-07
335	3562.I13.GZ43_507815	M26049	Yeast (S.cerevisiae) RAD9 protein (required for cell cycle arrest during DNA repair) gene, complete cds	4.00E-06
336	3562.I15.GZ43_507847	AF310880	Barbatula barbatula microsatellite Bbar5 sequence	1.60E-07

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
337	3562.J09.GZ43_507752	AF236642	Calothrix parietina clone 102-2A 16S-23S internal transcribed spacer, complete sequence, and tRNA-Ile and tRNA-Ala genes, complete sequence	3.30E-07
338	3562.J13.GZ43_507816	AC010728	Homo sapiens BAC clone RP11-258E22 from Y, complete sequence	1.30E-05
339	3562.K04.GZ43_507673	S79777	{specific DNA probe for Plasmodium vivax pARC 1153} [Plasmodium vivax, host=human, Genomic, 665 nt]	5.40E-07
340	3562.K08.GZ43_507737	AJ403240	M.musculus DNA for vimentin-binding fragment VimB8	2.00E-06
341	3562.L12.GZ43_507802	AE007840	Clostridium acetobutylicum ATCC824 section 328 of 356 of the complete genome	5.80E-07
342	3562.N24.GZ43_507996	AF255609	Homo sapiens high mobility group protein HMG1 gene, exons 1 and 2, partial cds	2.00E-07
343	3562.O11.GZ43_507789	M15027	Human myelin proteolipid protein gene, exon 2	1.00E-06
344	3562.O18.GZ43_507901	AL050208	Homo sapiens mRNA; cDNA DKFZp586F2323 (from clone DKFZp586F2323)	2.40E-07
345	3562.O20.GZ43_507933	AY020756	Oryza sativa microsatellite MRG3081 containing (TA) <sub>13</sub> , genomic sequence	4.90E-08
346	3562.P21.GZ43_507950	AF036318	Skeletonema costatum cyclin (CYCL) gene, partial cds	7.20E-07
347	3562.P23.GZ43_507982	AF126719	Plasmodium falciparum cAMP-dependent protein kinase (pka) gene, complete cds	3.00E-06
348	3565.A23.GZ43_508351	AL122065	Homo sapiens mRNA; cDNA DKFZp434N011 (from clone DKFZp434N011)	1.50E-07
349	3565.B05.GZ43_508064	AF163325	Trichoderma harzianum mitochondrial plasmid pThr1, complete plasmid sequence	1.50E-07
350	3565.B13.GZ43_508192	X62689	T.retusa DNA for brachiopod cubitus-interruptus dominant (ciD) homologue	9.00E-06
351	3565.B14.GZ43_508208	M29929	Human insulin receptor (allele 1) gene, exons 14, 15, 16 and 17	4.30E-12
352	3565.C04.GZ43_508049	AE006183	Pasteurella multocida PM70 section 150 of 204 of the complete genome	2.00E-06
353	3565.C06.GZ43_508081	S79836	SCPx/SCP2=sterol carrier protein x/sterol carrier protein 2 (promoter) [human, Genomic, 3575 nt]	3.00E-06
354	3565.C17.GZ43_508257	L13937	Bovine phospholipase C mRNA, complete cds	3.00E-07
355	3565.D14.GZ43_508210	M37818	Human keratin (psi-K-alpha) pseudogene, exons 4,5,6,7 and 8, and keratin (psi-K-beta) pseudogene, complete cds	3.50E-08
356	3565.D17.GZ43_508258	Z19005	C.pasteurianum gene for ferredoxin	1.00E-06

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
357	3565.D19.GZ43_508290	AE007758	<i>Clostridium acetobutylicum</i> ATCC824 section 246 of 356 of the complete genome	3.00E-06
358	3565.E16.GZ43_508243	L42813	<i>Protopterus dolloi</i> complete mitochondrial genome	2.49E-04
359	3565.G07.GZ43_508101	U97500	<i>Homo sapiens</i> butyrophilin (BT3.3) gene, exons 1-4	1.30E-05
360	3565.G09.GZ43_508133	M95098	<i>Bos taurus</i> lysozyme gene (cow 2), complete cds	1.26E-04
361	3565.G22.GZ43_508341	AJ400873	<i>Homo sapiens</i> partial GPLD1 gene for glycosylphosphatidylinositol phospholipase D, exons 15-20	1.40E-09
362	3565.H06.GZ43_508086	U67465	<i>Methanococcus jannaschii</i> section 7 of 150 of the complete genome	6.10E-07
363	3565.H10.GZ43_508150	M15350	<i>Bacillus</i> sp. strain 170 beta-lactamase gene, complete cds	5.70E-08
364	3565.H11.GZ43_508166	AB044878	<i>Equus caballus</i> DNA, microsatellite TKY378	3.20E-09
365	3565.H15.GZ43_508230	AL122122	<i>Homo sapiens</i> mRNA; cDNA DKFZp434L098 (from clone DKFZp434L098)	5.00E-06
366	3565.H23.GZ43_508358	J05492	<i>E. coli</i> cytochrome O ubiquinol oxidase (cyoA, cyoB, cyoC, cyoD and cyoE genes, complete cds	1.00E-06
367	3565.H24.GZ43_508374	AE001417	<i>Plasmodium falciparum</i> chromosome 2, section 54 of 73 of the complete sequence	1.70E-10
368	3565.K15.GZ43_508233	AB062985	<i>Macaca fascicularis</i> brain cDNA clone:QmoA-10670, full insert sequence	6.90E-105
369	3565.L22.GZ43_508346	L81801	<i>Homo sapiens</i> (subclone 1_a2 from P1 H31) DNA sequences, complete sequence	1.30E-05
370	3565.M15.GZ43_508235	X08038	<i>Methanobacterium thermoautotrophicum</i> rpoT, rpoU, rpoV and rpoX genes for RNA polymerase subunits A, B', B" and C	1.10E-05
371	3565.M20.GZ43_508315	Z93381	<i>Caenorhabditis elegans</i> cosmid F28G4, complete sequence	1.20E-05
372	3565.N12.GZ43_508188	M21573	<i>Salmon (S.salar)</i> growth hormone gene, complete cds	5.70E-05
373	3565.N13.GZ43_508204	AK001163	<i>Homo sapiens</i> cDNA FLJ10301 fis, clone NT2RM2000032	5.20E-08
374	3565.N19.GZ43_508300	AF321321	<i>Homo sapiens</i> dopamine transporter (SLC6A3) gene, exon 15 and complete cds	2.00E-06
375	3565.O02.GZ43_508029	X59773	<i>Pisum sativum</i> mRNA for P protein, a part of glycine cleavage complex	1.30E-05
376	3565.O03.GZ43_508045	Z27113	<i>H. Sapiens</i> gene for RNA polymerase II subunit 14.4 kD	2.00E-15
377	3565.O07.GZ43_508109	X96607	<i>M.musculus</i> IgH 3' alpha enhancer DNA	6.40E-05
378	3565.O15.GZ43_508237	Z35484	<i>Thermoanaerobacter</i> sp. ATCC53627 cgtA gene	3.00E-06

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Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
379	3565.P03.GZ43_508046	U11292	Human Ki nuclear autoantigen mRNA, complete cds	6.40E-07
380	3565.P09.GZ43_508142	X56261	Yeast PPH1 gene for protein phosphatase 2A	1.00E-06
381	3565.P22.GZ43_508350	AE007790	Clostridium acetobutylicum ATCC824 section 278 of 356 of the complete genome	3.00E-06
382	3565.P24.GZ43_508382	X61146	N.tabacum NTP303 pollen specific mRNA	2.70E-05
383	3568.A10.GZ43_508545	U46925	Arabidopsis thaliana GTP-binding protein ATGB2 mRNA, complete cds	3.00E-06
384	3568.B02.GZ43_508418	U83640	Mus caroli Sp100 gene, exons 3 and 4	1.90E-08
385	3568.B05.GZ43_508466	BC008293	Homo sapiens, Similar to RIKEN cDNA A430101B06 gene, clone MGC:13017 IMAGE:3537789, mRNA, complete cds	3.20E-16
386	3568.C22.GZ43_508739	AF280797	Homo sapiens NPC-related protein NAG73 mRNA, complete cds	1.00E-06
387	3568.D23.GZ43_508756	AK022922	Homo sapiens cDNA FLJ12860 fis, clone NT2RP2003559	8.00E-06
388	3568.E17.GZ43_508661	AF068294	Homo sapiens HDCMB45P mRNA, partial cds	5.30E-09
389	3568.E20.GZ43_508709	AE006417	Lactococcus lactis subsp. lactis IL1403 section 179 of 218 of the complete genome	1.10E-05
390	3568.F06.GZ43_508486	U52198	Vibrio anguillarum flagellin E (flaE), flagellin D (flaD), and flagellin B (flaB) genes, complete cds, and (flaG) gene, partial cds	2.20E-05
391	3568.F07.GZ43_508502	Z23599	H. sapiens (D135263) DNA segment containing (CA) repeat; clone AFM210yg11; single read	1.90E-08
392	3568.F11.GZ43_508566	AE007525	Clostridium acetobutylicum ATCC824 section 13 of 356 of the complete genome	4.20E-07
393	3568.F12.GZ43_508582	D50416	Mouse mRNA for AREC3, complete cds	1.90E-05
394	3568.F22.GZ43_508742	AF025900	Histronicus histronicus CA dinucleotide repeat locus Hhmicro1	7.80E-07
395	3568.G10.GZ43_508551	U66074	Trichomonas foetus putative superoxide dismutase 2 (SOD2) gene, complete cds	9.70E-07
396	3568.G12.GZ43_508583	AB062941	Macaca fascicularis brain cDNA clone:QfIA-14927, full insert sequence	9.50E-47
397	3568.G24.GZ43_508775	L27221	Giardia intestinalis pyruvate:flavodoxin oxidoreductase and flanking genes	3.20E-05
398	3568.H20.GZ43_508712	X75887	B. taurus Brevican mRNA	4.70E-05
399	3568.J10.GZ43_508554	AF194829	Tetragonia tetragonioides NADH dehydrogenase (ndhF) gene, partial cds; chloroplast gene for chloroplast product	2.00E-06
400	3568.J22.GZ43_508746	Y11031	C. coli pldA gene	1.00E-06
401	3568.K01.GZ43_508411	AL137751	Homo sapiens mRNA; cDNA DKFZp434I0812 (from clone DKFZp434I0812); partial cds	3.00E-06

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
402	3568.K04.GZ43_508459	Z82295	R.prowazekii genomic DNA fragment (clone A153F)	7.20E-08
403	3568.L04.GZ43_508460	AL050105	Homo sapiens mRNA; cDNA DKFZp586H0519 (from clone DKFZp586H0519); partial cds	1.00E-05
404	3568.M03.GZ43_508445	L76259	Homo sapiens PTS gene, complete cds	8.00E-06
405	3568.M13.GZ43_508605	X61218	M.musculus cervicolor (strain CRP) Tcp-1 gene for t-complex polypeptide 1, exons 8-10	3.10E-09
406	3568.N11.GZ43_508574	AL079296	Homo sapiens mRNA full length insert cDNA clone EUOIMAGE 609395	2.00E-06
407	3568.O17.GZ43_508671	AF078848	Homo sapiens BUP mRNA, complete cds	9.50E-09
408	3568.P04.GZ43_508464	AB041548	Mus musculus brain cDNA, clone MNCb-3816, similar to AF171875 g1-related zinc finger protein (Mus musculus)	5.00E-06
409	3568.P18.GZ43_508688	AL358951	Human DNA sequence from clone RP3-456L16 on chromosome 6, complete sequence [Homo sapiens]	3.00E-07
410	3568.P19.GZ43_508704	U43542	Nicotiana tabacum diphenol oxidase mRNA, complete cds	2.00E-06
411	3571.A04.GZ43_508833	AF017116	Homo sapiens type-2 phosphatidic acid phosphohydrolase (PAP2) mRNA, complete cds	2.40E-07
412	3571.A07.GZ43_508881	L81867	Homo sapiens (subclone 1_a8 from P1 H54) DNA sequence, complete sequence	9.00E-06
413	3571.A08.GZ43_508897	X85041	H.sapiens PE5L gene ALU repeat region	2.00E-06
414	3571.A11.GZ43_508945	U19361	Petromyzon marinus neurofilament subunit NF-180 mRNA, complete cds	4.70E-08
415	3571.A14.GZ43_508995	AL022342	Human DNA sequence from clone RP1-29M10 on chromosome 20, complete sequence [Homo sapiens]	7.00E-05
416	3571.A22.GZ43_509121	U09448	Vaucheria bursata protein synthesis elongation factor Tu (tufA) gene, chloroplast gene encoding chloroplast protein, partial cds	7.20E-07
417	3571.B13.GZ43_508978	AE002555	Neisseria meningitidis serogroup B strain MC58 section 197 of 206 of the complete genome	4.40E-05
418	3571.B22.GZ43_509122	AE002555	Neisseria meningitidis serogroup B strain MC58 section 197 of 206 of the complete genome	4.50E-05
419	3571.C08.GZ43_508899	AJ010154	Saguinus oedipus msp-E1 gene	1.10E-17
420	3571.D04.GZ43_508836	AF125460	Cacenorhabditis elegans cosmid Y9D1A	3.60E-07
421	3571.D07.GZ43_508884	U51654	Barbus barbus x Barbus meridionalis microsatellite clone no.37	8.72E-02
422	3571.E02.GZ43_508805	AF329081	Bos taurus AMP-activated protein kinase gamma-1 (PRKAG1) gene, partial cds	4.40E-33
423	3571.E10.GZ43_508933	M96068	Madagascar periwinkle hydroxymethylglutaryl-CoA reductase (HMGR) mRNA, complete cds	3.30E-08



Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
424	3571.E16.GZ43_509029	AE006429	<i>Lactococcus lactis</i> subsp. <i>lactis</i> IL 1403 section 191 of 218 of the complete genome	1.30E-05
425	3571.F06.GZ43_508870	AL137296	Homo sapiens mRNA; cDNA DKFZp434M0416 (from clone DKFZp434M0416)	4.40E-07
426	3571.F16.GZ43_509030	M58478	Human cystic fibrosis transmembrane conductance regulator gene, 5' end	6.30E-05
427	3571.F23.GZ43_509142	AF038397	Mus musculus glutaminase (Gls) gene, partial 3' sequence	4.70E-08
428	3571.G22.GZ43_509127	M80596	Saccharomyces cerevisiae VAC1 gene (required for vacuole inheritance and vacuole protein sorting), complete cds	7.00E-06
429	3571.G24.GZ43_509159	Z75330	H.sapiens mRNA for nuclear protein SA-1	1.00E-46
430	3571.H01.GZ43_508792	U71144	Influenza A virus H3N2 A/Akita/1/94 nucleoprotein (NP) gene, complete cds	1.90E-05
431	3571.H10.GZ43_508936	AF038564	Homo sapiens atrophin-1 interacting protein 4 (AIP4) mRNA, partial cds	6.60E-53
432	3571.H12.GZ43_508968	K00131	mouse b2 repeat sequence from clone mm61	3.00E-08
433	3571.H16.GZ43_509032	AF179564	Homo sapiens GTF2I-like sequence within duplicated segment of Williams syndrome region	1.20E-23
434	3571.H18.GZ43_509064	AE000331	Escherichia coli K12 MG1655 section 221 of 400 of the complete genome	1.45E-04
435	3571.I11.GZ43_508953	U20661	Dictyostelium discoideum unknown internal repeat protein gene, complete cds, and unknown orf1, orf2 and orf3 genes, partial cds	9.00E-06
436	3571.J07.GZ43_508890	M58478	Human cystic fibrosis transmembrane conductance regulator gene, 5' end	6.40E-05
437	3571.J08.GZ43_508906	AK021312	Mus musculus 13 days embryo stomach cDNA, RIKEN full-length enriched library, clone:D530039A21, full insert sequence	3.60E-08
438	3571.J09.GZ43_508922	X66483	D.discoideum gp80 gene	8.90E-07
439	3571.J14.GZ43_509002	L77119	Methanococcus jannaschii small extra-chromosomal element, complete sequence.~	1.40E-05
440	3571.L01.GZ43_508796	AK005500	Mus musculus adult female placenta cDNA, RIKEN full-length enriched library, clone:1600019O04, full insert sequence	6.00E-06
441	3571.M17.GZ43_509053	AF085681	Mus musculus tubby like protein 1 (Tulp1) mRNA, complete cds	5.00E-06
442	3571.M19.GZ43_509085	D10487	B.thermoglucosidase gene for oligo-1,6-glucosidase	9.00E-06
443	3571.M24.GZ43_509165	M97680	Bluetongue virus type 2 genomic RNA sequence	2.00E-06
444	3571.N09.GZ43_508926	X86100	R.norvegicus BSP gene	3.40E-07

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
445	3571.N14.GZ43_509006	D32007	Mouse mRNA for a homologue of human CBFAT21(Mtg8a), complete cds	1.20E-08
446	3571.N17.GZ43_509054	Z68755	Human DNA sequence from cosmid L118D5, Huntington's Disease Region, chromosome 4p16.3	1.70E-10
447	3571.N22.GZ43_509134	D00326	Porcine rotavirus (strain Gottfried), VP6 gene, complete cds	1.00E-06
448	3571.O08.GZ43_508911	X66483	D.discoideum gp80 gene	8.20E-07
449	3574.A20.GZ43_509473	AJ271814	Drosophila melanogaster mRNA for meso18E protein	1.70E-07
450	3574.B01.GZ43_509170	U93261	Homo sapiens DESP4P1 pseudogene sequence	1.00E-06
451	3574.B04.GZ43_509218	Y08207	C.elaphus mitochondrial tRNA-Thr, tRNA-Pro and tRNA-Phe genes	1.40E-14
452	3574.B10.GZ43_509314	AL161991	Homo sapiens mRNA; cDNA DKFZp761C169 (from clone DKFZp761C169); partial cds	3.00E-06
453	3574.B14.GZ43_509378	D79208	Apis mellifera mRNA for alpha-glucosidase, complete cds	7.00E-06
454	3574.B24.GZ43_509538	AE007758	Clostridium acetobutylicum ATCC824 section 246 of 356 of the complete genome	3.00E-06
455	3574.C09.GZ43_509299	AF057708	Populus balsamifera subsp. trichocarpa PTD protein (PTD) gene, complete cds	2.40E-07
456	3574.C10.GZ43_509315	AE005602	Bscherichia coli O157:H7 EDL933 genome, contig 3 of 3, section 221 of 290	9.70E-05
457	3574.C12.GZ43_509347	AJ223633	Enterococcus faecium genes encoding enterocin L50A and enterocin L50B plus 5' and 3' flanking regions	9.50E-07
458	3574.C14.GZ43_509379	X99710	L.lactis ORF, genes homologous to vsf-1 and pepF2 and gene encoding protein homologous to methyltransferase	4.00E-06
459	3574.C16.GZ43_509411	AF092920	Chlorohydra viridissima head-activator binding protein precursor (HAB) mRNA, complete cds	3.00E-07
460	3574.C23.GZ43_509523	AB047856	Oryza sativa Ub-CEP52-2 gene for ubiquitin fused to ribosomal protein L40, complete cds	5.00E-08
461	3574.D02.GZ43_509188	AB060225	Macaca fascicularis brain cDNA clone: QfLA-14955, full insert sequence	5.70E-07
462	3574.D12.GZ43_509348	M58478	Human cystic fibrosis transmembrane conductance regulator gene, 5' end	6.00E-05
463	3574.E02.GZ43_509189	L37347	Human integral membrane protein (Nramp2) mRNA, partial	2.00E-06
464	3574.E03.GZ43_509205	X05817	Bovine papillomavirus type 4 (BPV-4) genome	6.00E-06
465	3574.E14.GZ43_509381	U67507	Methanococcus jaunaschii section 49 of 150 of the complete genome	3.40E-05
466	3574.F10.GZ43_509318	M24376	Mouse zinc finger protein (krox-20) gene, exon 1	3.80E-08

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
467	3574.F18.GZ43_509446	AF184170	Sparus aurata elongation factor 1-alpha (EF1-alpha) mRNA, complete cds	3.40E-07
468	3574.F23.GZ43_509526	Z29486	R.norvegicus (Sprague Dawley) mRNA for AMP-activated protein kinase	9.00E-06
469	3574.G07.GZ43_509271	AF064079	Plasmodium gallinaceum endochitinase precursor, mRNA, complete cds	1.40E-07
470	3574.G11.GZ43_509335	AF032872	Rattus norvegicus potassium channel regulatory protein KChAP mRNA, complete cds	7.40E-07
471	3574.H07.GZ43_509272	J04718	Human proliferating cell nuclear antigen (PCNA) gene, complete cds	3.10E-07
472	3574.I02.GZ43_509193	AF200361	Rattus norvegicus cytochrome P450 4F1 (Cyp4F1) gene, complete cds	1.50E-05
473	3574.I07.GZ43_509273	M29688	S.cerevisiae PMS1 gene encoding DNA mismatch repair protein, complete cds	1.20E-08
474	3574.J11.GZ43_509338	Z24104	H. sapiens (D12S338) DNA segment containing (CA) repeat; clone AFM291wd9; single read	3.20E-07
475	3574.J14.GZ43_509386	AB008430	Homo sapiens mRNA for CDEP, complete cds	4.70E-05
476	3574.J23.GZ43_509530	AP000384	Arabidopsis thaliana genomic DNA, chromosome 3, P1 clone:MCE21	7.10E-07
477	3574.K12.GZ43_509355	AB031814	Mus musculus oatp2 mRNA for organic anion transporting polypeptide 2, complete cds	1.50E-05
478	3574.K20.GZ43_509483	AF126719	Plasmodium falciparum cAMP-dependent protein kinase (pka) gene, complete cds	3.00E-06
479	3574.L07.GZ43_509276	U53400	Rattus norvegicus chromosome 10 microsatellite sequence D10Mc021	8.94E-02
480	3574.M03.GZ43_509213	AB000404	Rice grassy stunt virus genomic RNA6 for 20.6K major nonstructural protein and 36.4K protein, complete cds	5.60E-07
481	3574.M23.GZ43_509533	U18056	Lycopersicon esculentum 1-amino-cyclopropane-1-carboxylate synthase (LE-ACS1A) gene, complete cds	3.40E-07
482	3574.N04.GZ43_509230	L48479	Homo sapiens (subclone 6_h1 from P1 H21) DNA sequence	3.30E-09
483	3574.N10.GZ43_509326	M58150	Bovine lactoperoxidase (LPO) mRNA, complete cds	3.60E-05
484	3574.N12.GZ43_509358	AF182950	Homo sapiens HEX (HEX) gene, partial cds and 5' flanking sequence	9.00E-06
485	3574.N20.GZ43_509486	AB006904	Sulfolobus solfataricus section 263 of 272 of the complete genome	3.00E-06
486	3574.P07.GZ43_509280	U60232	Homo sapiens cysteine dioxygenase (CDO-1) gene, 5' flanking region and exons 1 and 2	6.30E-08
487	3574.P17.GZ43_509440	AC002218	Homo sapiens (subclone 2_c1 from P1 H43) DNA sequence, complete sequence	5.30E-08
488	3577.A06.GZ43_509633	U28328	Bos taurus dinucleotide repeat RM154, tandem repeat region	3.40E-27

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
489	3577.A18.GZ43_509825	X58774	Herpesvirus saimiri sRNA1, sRNA2, sRNA3 and sRNA4 genes for small viral RNAs	1.00E-06
490	3577.B12.GZ43_509730	BC008400	Homo sapiens, postmitotic segregation increased (S. cerevisiae) 2, clone IMAGE:4273792, mRNA	2.50E-05
491	3577.B15.GZ43_509778	M61127	Drosophila melanogaster GTP-binding protein (arf-like) gene, complete cds	1.10E-05
492	3577.B19.GZ43_509842	AF135526	Homo sapiens clone MINT26 colon cancer differentially methylated CpG island genomic sequence	1.00E-06
493	3577.E19.GZ43_509845	AF063864	Schizosaccharomyces pombe essential nuclear protein Mcm3p (mcm3+) gene, complete cds	1.00E-06
494	3577.F02.GZ43_509574	U37434	Danio rerio L-isoaspartate (D-aspartate) O-methyltransferase (PCMT) mRNA, complete cds	5.10E-08
495	3577.G07.GZ43_509655	AF001893	Human MEN1 region clone epsilon/beta mRNA, 3' fragment	3.00E-06
496	3577.G13.GZ43_509751	M83821	Xenopus laevis mucin B.1 consensus repeat mRNA	2.10E-07
497	3577.H06.GZ43_509640	AK007565	Mus musculus 10 day old male pancreas cDNA, RIKEN full-length enriched library, clone:1810020K22, full insert sequence	8.00E-07
498	3577.H08.GZ43_509672	L81912	Homo sapiens (subclone 2_g5 from PAC H74) DNA sequence, complete sequence	2.40E-07
499	3577.H18.GZ43_509832	AL157461	Homo sapiens mRNA; cDNA DKFZp434K152 (from clone DKFZp434K152)	4.00E-06
500	3577.I01.GZ43_509561	U35006	Carcharhinus plumbeus Ig lambda b1a light chain gene, complete cds	2.00E-06
501	3577.I17.GZ43_509817	AF157252	Gongronella butleri translation elongation factor 1-alpha (EF-1alpha) gene, partial cds	1.00E-06
502	3577.J04.GZ43_509610	AF338249	Sus scrofa thyroid-stimulating hormone receptor mRNA, complete cds	2.00E-06
503	3577.K06.GZ43_509643	AB000264	Bacillus firmus DNA for beta-amylase, partial cds	5.00E-07
504	3577.K14.GZ43_509771	X15441	Aspergillus nidulans mitochondrial ndhC and oxib genes for NADH dehydrogenase subunit 3 and cytochrome oxidase subunit II	1.00E-06
505	3577.K23.GZ43_509915	X52952	Rat mRNA for c-mos	3.00E-06
506	3577.L10.GZ43_509708	X60578	Hepatitis C genomic RNA for putative envelope protein (RE56 isolate)	3.70E-07
507	3577.N10.GZ43_509710	Z75121	S. cerevisiae chromosome XV reading frame ORF YOR213c	4.50E-09
508	3577.N14.GZ43_509774	M90058	Human serglycin gene, exons 1,2, and 3	5.00E-06
509	3577.O17.GZ43_509823	L19141	Lupinus albus L-asparaginase gene, complete cds	9.10E-08

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
510	3577.O22.GZ43_509903	AL031008	Human DNA sequence from clone 360A4 on chromosome 16. Contains ESTs, complete sequence [Homo sapiens]	5.60E-08
511	3577.P02.GZ43_509584	AK006176	Mus musculus adult male testis cDNA, RIKEN full-length enriched library, clone:1700020M10, full insert sequence	4.60E-08
512	3577.P07.GZ43_509664	U05822	Human proto-oncogene BCL3 gene, exon 2	2.40E-14
513	3577.P23.GZ43_509920	AJ010341	Homo sapiens PISSLRE gene, exons 1, 2, and 3 and joined CDS	1.00E-11
514	3580.A04.GZ43_509985	AJ010213	Mus musculus beta-dystrobrevin gene, exon 10	8.20E-07
515	3580.A09.GZ43_510065	AB037862	Homo sapiens mRNA for KIAA1441 protein, partial cds	6.30E-15
516	3580.A13.GZ43_510129	U17832	Symploce pallens mitochondrion 16S ribosomal RNA, partial sequence	7.80E-07
517	3580.A14.GZ43_510145	X89414	A.thaliana DNA for pyruvate-5-carboxylase synthetase gene	6.00E-06
518	3580.B01.GZ43_509938	U67487	Methanococcus jannaschii section 29 of 150 of the complete genome	9.00E-05
519	3580.C01.GZ43_509939	X14898	Hamster p7 preinsertion DNA	2.00E-06
520	3580.C03.GZ43_509971	X76302	H.sapiens RY-1 mRNA for putative nucleic acid binding protein	3.70E-07
521	3580.C05.GZ43_510003	Z22923	M.musculus alpha2 (IX) collagen gene, complete CDS	1.60E-05
522	3580.D07.GZ43_510036	AB062941	Macaca fascicularis brain cDNA clone:QfIA-14927, full insert sequence	9.80E-22
523	3580.D22.GZ43_510276	M84136	Flaveria chloraefolia flavonol 4'-sulfotransferase mRNA, complete cds	4.00E-06
524	3580.E02.GZ43_509957	AE001002	Archaeoglobus fulgidus section 105 of 172 of the complete genome	3.90E-05
525	3580.E08.GZ43_510053	U48431	Drosophila pseudoobscura alpha-amylase (Amy3) pseudogene, complete cds	3.00E-06
526	3580.E10.GZ43_510085	Z64717	H.sapiens CpG island DNA genomic MseI fragment, clone 161e9, forward read cpg161e9.ft1a	9.60E-19
527	3580.E19.GZ43_510229	M64984	Candida tropicalis open reading frame DNA sequence	2.00E-06
528	3580.E21.GZ43_510261	M84136	Flaveria chloraefolia flavonol 4'-sulfotransferase mRNA, complete cds	5.00E-06
529	3580.E23.GZ43_510293	AB033570	Epilatreus burgeri hgpTPR5a mRNA, partial cds	2.00E-06
530	3580.G03.GZ43_509975	Y14277	Drosophila melanogaster mRNA for nuclear protein SA	1.10E-05
531	3580.G13.GZ43_510135	AK018491	Mus musculus adult male colon cDNA, RIKEN full-length enriched library, clone:9030408N04, full insert sequence	4.40E-08
532	3580.G14.GZ43_510151	AF142660	Lama glama microsatellite LCA90 sequence	2.60E-07

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
533	3580.G18.GZ43_510215	D86226	<i>Spinacia oleracea</i> DNA for nitrate reductase, complete cds	2.60E-05
534	3580.G19.GZ43_510231	U60502	Glycine max actin (Soy119) gene, partial cds	7.00E-06
535	3580.G20.GZ43_510247	D38524	Human mRNA for 5'-nucleotidase	4.80E-11
536	3580.G24.GZ43_510311	AF084480	<i>Mus musculus</i> Williams-Beuren syndrome deletion transcript 9 homolog (Wbscr9) mRNA, complete cds	5.00E-06
537	3580.H12.GZ43_510120	X78423	<i>D. carota</i> (Queen Anne's Lace) Inv*De3 gene, 4444bp	4.00E-06
538	3580.H16.GZ43_510184	Y13786	Homo sapiens mRNA for meltrin-beta/ADAM 19 homologue	4.50E-10
539	3580.H22.GZ43_510280	X62578	<i>C. caldarium</i> plastid genes ompR, psbD, psbC, rps16 and groEL	2.50E-05
540	3580.I06.GZ43_510025	X51344	<i>Spiroplasma virus</i> (SpV1-R8A2 B) complete genome	4.70E-07
541	3580.I08.GZ43_510057	X02761	Human mRNA for fibronectin (FN precursor)	1.02E-04
542	3580.I18.GZ43_510217	BC007856	Homo sapiens, clone MGC:14337 IMAGE:4298428, mRNA, complete cds	2.60E-10
543	3580.J10.GZ43_510090	AF068206	<i>Rangifer tarandus</i> microsatellite NVHRT16 sequence	4.40E-11
544	3580.J12.GZ43_510122	AE008323	<i>Agrobacterium tumefaciens</i> strain C58 linear chromosome, section 127 of 187 of the complete sequence	9.30E-05
545	3580.J18.GZ43_510218	AF222689	Homo sapiens protein arginine N-methyltransferase 1 (HRMT1L2) gene, complete cds, alternatively spliced	1.50E-05
546	3580.J20.GZ43_510250	M31651	Homo sapiens sex hormone-binding globulin (SHBG) gene, complete cds	3.80E-07
547	3580.J21.GZ43_510266	AB054062	<i>Pagrus major</i> lpl mRNA for lipoprotein lipase, complete cds	3.00E-06
548	3580.K03.GZ43_509979	AE007607	<i>Clostridium acetobutylicum</i> ATCC824 section 95 of 356 of the complete genome	5.00E-05
549	3580.K05.GZ43_510011	Z15027	H.sapiens HLA class III DNA	3.70E-08
550	3580.K21.GZ43_510267	AF135826	<i>Mus musculus</i> neuronal nitric oxide synthase (NOS-1) gene, exon 1c and 5'-flanking sequence	2.20E-09
551	3580.L09.GZ43_510076	AL049333	Homo sapiens mRNA; cDNA DKFZp564M116 (from clone DKFZp564M116)	3.40E-13
552	3580.L10.GZ43_510092	AF278587	<i>Borrelia burgdorferi</i> strain BC-1 outer surface protein C (ospC) gene, partial cds	2.00E-06
553	3580.L12.GZ43_510124	D14664	Human mRNA for KIAA0022 gene, complete cds	1.10E-05
554	3580.L13.GZ43_510140	K02269	Human ERV3 (endogenous retrovirus 3) gag gene	3.30E-07
555	3580.L17.GZ43_510204	U60232	Homo sapiens cysteine dioxygenase (CDO-1) gene, 5' flanking region and exons 1 and 2	2.00E-07

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
556	3580.M01.GZ43_50994 9	U53400	Rattus norvegicus chromosome 10 microsatellite sequence D10Mco21	4.54E-01
557	3580.M16.GZ43_51018 9	AE006406	Lactococcus lactis subsp. lactis IL1403 section 168 of 218 of the complete genome	3.00E-06
558	3580.M17.GZ43_51020 5	AF348584	Arabidopsis thaliana unknown protein (T8K14.7) mRNA, complete cds	6.70E-07
559	3580.M18.GZ43_51022 1	X69908	H.sapiens gene for mitochondrial ATP synthase c subunit (P2 form)	1.00E-05
560	3580.M23.GZ43_51030 1	M17326	Mouse endogenous murine leukemia virus polytropic provirus DNA, complete cds	9.00E-06
561	3580.N10.GZ43_510094	AF103970	Lasioglossum rohweri cytochrome oxidase I (COI) gene, mitochondrial gene encoding mitochondrial protein, partial cds	1.00E-06
562	3580.N11.GZ43_510110	Z80362	H.sapiens HLA-DRB pseudogene, exon 1; Xenopus laevis XNLR-1 mRNA, complete cds	6.10E-11
563	3580.N14.GZ43_510158	AB014462	Anomochloa marantioidea maturase (matK) gene, complete cds; chloroplast gene for chloroplast product	1.60E-05
564	3580.N15.GZ43_510174	AF164381	Macaca fascicularis brain cDNA, clone:OmpA-14303	1.00E-06
565	3580.N23.GZ43_510302	AB047880	H. aspersa cytoplasmic intermediate filament gene exons 2 to 6	2.00E-06
566	3580.O02.GZ43_509967	X55948	Homo sapiens platelet/endothelial cell adhesion molecule-1 (PECAM-1) gene, exon 14	4.00E-06
567	3580.O06.GZ43_510031	L34649	H.sapiens mig-5 gene	4.00E-06
568	3580.O07.GZ43_510047	Z30183	H.sapiens ribosomal protein L11 gene, complete cds	3.00E-05
569	3580.O08.GZ43_510063	AF101385	Homo sapiens BAC clone RP11-221K4 from Y, complete sequence	1.80E-08
570	3580.P04.GZ43_510000	AC016707	Thermotoga neapolitana galactose utilization operon, complete sequence	1.80E-08
571	3580.P05.GZ43_510016	AF055482	Rattus norvegicus CD94 (Cd94) mRNA, complete cds	8.00E-07
572	3580.P14.GZ43_510160	AF009133	Human papillomavirus type 80 E6, E7, E1, E2, E4, L2, and L1 genes	7.50E-08
573	3580.P19.GZ43_510240	Y15176	Chlamydomonas moewusii chloroplast DNA for ORF 563 and transfer RNA-Thr	7.00E-06
574	3583.B06.GZ43_510402	X51398	Hexachacta amabilis 16S ribosomal RNA gene, mitochondrial gene encoding mitochondrial RNA, partial sequence	3.00E-06
575	3583.B07.GZ43_510418	U39382	erythropoietin receptor [human, placental, Genomic, 8647 nt]	5.50E-08
576	3583.B10.GZ43_510466	S45332	Caenorhabditis elegans clone CS2E2, complete sequence	3.90E-10
577	3583.B11.GZ43_510482	AC006623		4.00E-06

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
578	3583.D15.GZ43_510548	AF242297	Homo sapiens phosphatidylinositol 3-kinase, promoter and exon 1	3.80E-08
579	3583.D22.GZ43_510660	Z23548	H. sapiens (D105540) DNA segment containing (CA) repeat; clone AFM205xe11; single read	3.20E-07
580	3583.E11.GZ43_510485	X69737	E. coli chloroplast rbcL gene for ribulose-1,5-bisphosphate-carboxylase and promoter region	1.30E-08
581	3583.E13.GZ43_510517	AB007856	Homo sapiens KIAA0396 mRNA, partial cds	2.20E-05
582	3583.E15.GZ43_510549	X74131	H. nelsoni small subunit ribosomal RNA	7.00E-06
583	3583.E17.GZ43_510581	AE006633	Streptococcus pyogenes M1 GAS strain SF370, section 162 of 167 of the complete genome	2.40E-07
584	3583.F24.GZ43_510694	J02846	Human tissue factor gene, complete cds	7.40E-07
585	3583.G09.GZ43_510455	X88789	P. sativum mRNA for starch synthase (2035 bp)	2.10E-05
586	3583.G16.GZ43_510567	AK000735	Homo sapiens cDNA FLJ20728 fis, clone HEP11763	4.70E-07
587	3583.G17.GZ43_510583	AK026822	Homo sapiens cDNA: FLJ23169 fis, clone LNG09957	2.60E-05
588	3583.G21.GZ43_510647	U13044	Human nuclear respiratory factor-2 subunit alpha mRNA, complete cds	2.00E-06
589	3583.H03.GZ43_510360	M26222	African green monkey origin of replication (ORS9) region	1.00E-13
590	3583.H12.GZ43_510504	X01669	Human c-k-ras oncogene exon 2 from lung carcinoma pr310	3.20E-08
591	3583.H13.GZ43_510520	AK022380	Homo sapiens cDNA FLJ12318 fis, clone MAMMA1002068	2.00E-06
592	3583.H15.GZ43_510552	L77119	Methanococcus jannaschii small extra-chromosomal element, complete sequence...	1.60E-05
593	3583.J02.GZ43_510346	AJ007302	Sus scrofa triadin gene	1.00E-06
594	3583.K08.GZ43_510443	D63902	Mouse mRNA for estrogen-responsive finger protein, complete cds	2.50E-11
595	3583.K10.GZ43_510475	U11816	Lactobacillus strain 30A ornithine decarboxylase (odc) gene, complete cds	1.00E-05
596	3583.K11.GZ43_510491	X73416	W. suaveolens mitochondrialorf1	6.00E-06
597	3583.K14.GZ43_510539	U04367	Bacillus thuringiensis dakota HD511 CryIII delta-endotoxin gene, partial cds	1.20E-05
598	3583.K17.GZ43_510587	AE004129	Vibrio cholerae chromosome I, section 37 of 251 of the complete chromosome	8.00E-06
599	3583.K23.GZ43_510683	AE001410	Plasmodium falciparum chromosome 2, section 47 of 73 of the complete sequence C.stercorarium celZ gene for endo-beta-1,4-glucanase (Avicelase I)	4.00E-06
600	3583.L05.GZ43_510396	X55299	Homo sapiens SOS1 (SOS1) gene, partial cds	1.00E-05
601	3583.L08.GZ43_510444	AF106953	Soybean chloroplast phytochrome A (phyA) gene, complete cds	7.50E-09
602	3583.L09.GZ43_510460	L34842		2.40E-05



Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
603	3583.L17.GZ43_510588	X65223	<i>Trubrum</i> mitochondrion genes for cytochrome oxidase I, cytochrome oxidase II, ATPase 9, NADH dehydrogenase subunit 4L, NADH dehydrogenase subunit 5, tRNA-Gln, tRNA-Met and tRNA-Arg	5.00E-06
604	3583.L21.GZ43_510652	AF106661	<i>Rattus norvegicus</i> glutathione S-transferase Yb4 (GstYb4) gene, complete cds	5.00E-06
605	3583.M08.GZ43_510445	BC005276	Homo sapiens, Similar to GR02 oncogene, clone IMAGE:4071652, mRNA	3.70E-07
606	3583.M10.GZ43_510477	Y00477	Human bone marrow serine protease gene (medullasin) (leukocyte neutrophil elastase gene)	4.70E-09
607	3583.M13.GZ43_510525	X73030	<i>S.cerevisiae</i> YGP1 gene	7.00E-06
608	3583.N09.GZ43_510462	AK018377	<i>Mus musculus</i> 16 days embryo lung cDNA, RIKEN full-length enriched library, clone:8430403M08, full insert sequence	4.60E-07
609	3583.O03.GZ43_510367	X72698	<i>P.pygmaeus</i> ZFY gene for Y-linked Zinc finger protein, final intron	3.00E-06
610	3583.O11.GZ43_510495	U40161	<i>Arabidopsis thaliana</i> type 2A protein serine/threonine phosphatase 55 kDa B regulatory subunit mRNA, complete cds	2.00E-06
611	3583.O17.GZ43_510591	U67567	<i>Methanococcus januszhii</i> section 109 of 150 of the complete genome	2.00E-06
612	3583.P09.GZ43_510464	AK021312	<i>Mus musculus</i> 13 days embryo stomach cDNA, RIKEN full-length enriched library, clone:D530039A21, full insert sequence	3.60E-08
613	3583.P19.GZ43_510624	U12920	<i>Caenorhabditis elegans</i> sex determination (tra-3) gene, exons 2-6	1.60E-05
614	3583.P22.GZ43_510672	AJ133800	Homo sapiens CPNE7 gene (partial), exon 2	7.60E-07
615	3590.A12.GZ43_512274	AF185661	<i>Glomus intraradices</i> strain FL208 18S ribosomal RNA, partial sequence; internal transcribed spacer 1, 5.8S ribosomal RNA and internal transcribed spacer 2, complete sequence; 26S ribosomal RNA, partial sequence	2.00E-06
616	3590.B01.GZ43_512099	M96068	Madagascar periwinkle hydroxymethylglutaryl-CoA reductase (HMGCR) mRNA, complete cds	7.40E-09
617	3590.B16.GZ43_512339	V01527	Mouse gene coding for major histocompatibility antigen. This is a class II antigen, I-A-beta	2.40E-12
618	3590.B21.GZ43_512419	AB028983	Homo sapiens mRNA for KIAA1060 protein, partial cds	1.70E-05
619	3590.C20.GZ43_512404	D86566	Human DNA for NOTCH4, partial cds	3.20E-07

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
620	3590.D03.GZ43_512133	D10371	Phocine distemper virus (PDV) genomic RNA for N, P, V, C, M, F, H and L protein	2.90E-05
621	3590.D19.GZ43_512389	M96163	Mus musculus (clone 2) serum inducible kinase (SNK) mRNA, mRNA sequence	7.80E-10
622	3590.D23.GZ43_512453	AF086485	Homo sapiens full length insert cDNA clone ZD93E02	7.70E-09
623	3590.E08.GZ43_512214	AF055278	Homo sapiens DNA repair protein XRCC4 (XRCC4) gene, exon 1	5.90E-12
624	3590.E10.GZ43_512246	AB001477	Helicobacter pylori, strain J99 section 38 of 132 of the complete genome	2.00E-06
625	3590.F01.GZ43_512103	AF080395	Entamoeba histolytica actin binding protein (abp2) mRNA, partial cds	2.00E-06
626	3590.F16.GZ43_512343	X79388	B.subtilis (168) prfA gene	1.20E-05
627	3590.G01.GZ43_512104	U32690	Haemophilus influenzae Rd section 5 of 163 of the complete genome	2.80E-05
628	3590.G02.GZ43_512120	U68040	Cochliobolus heterostrophus polyketide synthase (PKS1) gene, complete cds	1.25E-04
629	3590.H04.GZ43_512153	X66013	T.aestivum gene for cathepsin B (A116)	2.50E-07
630	3590.H06.GZ43_512185	X66177	M.musculus mRNA for Hox 2.7 protein	8.00E-06
631	3590.H09.GZ43_512233	AF012899	Sambucus nigra ribosome inactivating protein precursor mRNA, complete cds	3.40E-11
632	3590.H12.GZ43_512281	Y15724	Homo sapiens SERCA3 gene, exons 1-7 (and joined CDS)	2.00E-06
633	3590.H16.GZ43_512345	AF064079	Plasmodium gallinaceum endochitinase precursor, mRNA, complete cds	6.70E-09
634	3590.I16.GZ43_512346	L06280	Drosophila melanogaster adenine phosphoribosyltransferase (APRT) gene, complete cds	4.40E-07
635	3590.J01.GZ43_512107	X69573	T.reesei xyn1 gene, complete CDS	1.70E-07
636	3590.J02.GZ43_512123	AF092047	Homo sapiens homeobox protein Six3 (SIX3) gene, complete cds	4.00E-06
637	3590.J18.GZ43_512379	AB027966	Schizosaccharomyces pombe gene for Hypothetical protein, partial cds, clone:TB89	2.60E-08
638	3590.J21.GZ43_512427	AK014727	Mus musculus 0 day neonate head cDNA, RIKEN full-length enriched library, clone:4833419G08, full insert sequence	7.90E-08
639	3590.J22.GZ43_512443	AK020136	Mus musculus 12 days embryo male wolffian duct includes surrounding region cDNA, RIKEN full-length enriched library, clone:6720460K10, full insert sequence	5.90E-08
640	3590.K06.GZ43_512188	AF171890	Trimeresurus trigonocephalus cytochrome b (cytb) gene, partial cds; mitochondrial gene for mitochondrial product	3.00E-06
641	3590.K10.GZ43_512252	U16775	Human immunodeficiency virus type 1 isolate VEB6 reverse transcriptase (pol) gene, partial cds	6.00E-06

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Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
642	3590.K19.GZ43_512396	U40454	<i>Candida albicans</i> topoisomerase type I (CATOPI) gene, complete cds	3.00E-06
643	3590.L08.GZ43_512221	U52198	<i>Vibrio anguillarum</i> flagellin E (flaE), flagellin D (flaD), and flagellin B (flaB) genes, complete cds, and (flaG) gene, partial cds	2.00E-05
644	3590.L10.GZ43_512253	U01155	<i>Xenopus laevis</i> angiotensin II receptor mRNA, complete cds	4.00E-06
645	3590.M03.GZ43_512142	AF252499	<i>Bos taurus</i> clone MNB-88 microsatellite sequence	4.60E-08
646	3590.M04.GZ43_512158	AE007607	<i>Clostridium acetobutylicum</i> ATCC824 section 95 of 356 of the complete genome	4.50E-05
647	3590.M09.GZ43_512238	L04758	<i>Oryctolagus cuniculus</i> cytochrome P-450 (CYP4A4) gene, 5' end	1.00E-06
648	3590.N04.GZ43_512159	Z82038	<i>C.thermosaccharolyticum</i> etfB, etfA, hbd, thlA and actA genes	2.00E-06
649	3590.N19.GZ43_512399	U15603	<i>Saccharomyces cerevisiae</i> Csd3p (CSD3) gene, complete cds	4.00E-06
650	3590.N21.GZ43_512431	L19535	<i>Drosophila subobscura</i> sry alpha gene, complete cds	6.00E-06
651	3590.O08.GZ43_512224	L36588	<i>Homo sapiens</i> intron-encoded U22 small nucleolar RNA (UHG) gene	4.30E-07
652	3596.C02.GZ43_512500	L14849	<i>Drosophila melanogaster</i> cytoplasmic protein tyrosine phosphatase (PTP61F) mRNA, complete cds	8.90E-09
653	3596.C20.GZ43_512788	M60286	<i>Herpesvirus saimiri</i> immediate early region protein genes, complete cds	1.30E-07
654	3596.C22.GZ43_512820	X15121	Soybean Gyl gene for glycinin subunit G1	1.00E-06
655	3596.D01.GZ43_512485	Z78414	<i>Caenorhabditis elegans</i> cosmid W09D12, complete sequence	4.00E-06
656	3596.D07.GZ43_512581	M88242	Mouse glucocorticoid-regulated inflammatory prostaglandin G/H synthase (griPGHS) mRNA, complete cds	1.70E-05
657	3596.D09.GZ43_512613	X99710	<i>L.lactis</i> ORF, genes homologous to vsf-1 and pepF2 and gene encoding protein homologous to methyltransferase	5.00E-06
658	3596.D17.GZ43_512741	AF200361	<i>Rattus norvegicus</i> cytochrome P450 4F1 (Cyp4F1) gene, complete cds	1.40E-05
659	3596.E08.GZ43_512598	AF111848	<i>Homo sapiens</i> PRO0529 mRNA, complete cds	5.00E-06
660	3596.E22.GZ43_512822	X58178	<i>S.pyogenes</i> for emm41 gene	5.00E-06
661	3596.F10.GZ43_512631	AL390161	<i>Homo sapiens</i> mRNA; cDNA DKFZp761P0615 (from clone DKFZp761P0615)	2.00E-06
662	3596.G13.GZ43_512680	AJ000044	<i>Tenebrio molitor</i> LPCP29 gene	2.00E-06
663	3596.H04.GZ43_512537	U65018	<i>Dictyostelium discoideum</i> mannosyltransferase gene, complete cds	3.60E-07
664	3596.H10.GZ43_512633	AF104390	<i>Penaeus monodon</i> hyperglycemic hormone homolog PmSGP-V precursor; mRNA, complete cds	2.00E-06

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
665	3596.H17.GZ43_512745	D28915	Human gene for hepatitis C-associated microtubular aggregate protein p44, exon 9 and complete cds	1.00E-06
666	3596.H22.GZ43_512825	AF198250	Dictyostelium discoideum lim2 protein (limB) mRNA, complete cds	7.30E-07
667	3596.I06.GZ43_512570	U32444	Solanum lycopersicum phytochrome F (PHYF) gene, partial cds	1.10E-05
668	3596.I16.GZ43_512730	U32444	Solanum lycopersicum phytochrome F (PHYF) gene, partial cds	8.00E-06
669	3596.J04.GZ43_512539	D28596	Chicken gene for c-maf proto-oncogene product c-Maf, short form complete cds and long form 1st exon	9.30E-10
670	3596.J13.GZ43_512683	AB007856	Homo sapiens KIAA0396 mRNA, partial cds	2.40E-05
671	3596.K14.GZ43_512700	AC024752	Caenorhabditis elegans cosmid Y1B5A, complete sequence	3.00E-06
672	3596.K15.GZ43_512716	Y00469	Yeast mRNA for profilin	2.00E-06
673	3596.L01.GZ43_512493	X79703	O.aries gene for beta-casein	4.00E-06
674	3596.L08.GZ43_512605	AJ007313	Streptomyces coelicolor sigT, trxB and trxA genes, and ORF1 and ORF2	9.80E-07
675	3596.L13.GZ43_512685	AK018239	Mus musculus adult male medulla oblongata cDNA, RIKEN full-length enriched library, clone:6330563C09, full insert sequence	1.00E-06
676	3596.N02.GZ43_512511	AE001387	Plasmodium falciparum chromosome 2, section 24 of 73 of the complete sequence	1.00E-06
677	3596.N12.GZ43_512671	Z12841	O.uniculus mRNA for phospholipase	4.00E-06
678	3596.N15.GZ43_512719	U14186	Bos taurus general vesicular transport factor p115 mRNA, complete cds	1.70E-05
679	3596.N16.GZ43_512735	U41106	Caenorhabditis elegans cosmid W06A11	1.10E-05
680	3596.N21.GZ43_512815	AF097717	Homo sapiens 3'-phosphoadenosine 5'-phosphosulfate synthetase (PAPSS), exon 8	1.40E-07
681	3596.O10.GZ43_512640	AE001649	Chlamydia pneumoniae section 65 of 103 of the complete genome	1.10E-05
682	3596.O12.GZ43_512672	AC006623	Caenorhabditis elegans clone C52E2, complete sequence	4.00E-06
683	3596.P03.GZ43_512529	X82317	C.thummi CpY gene	1.49E-03
684	3596.P04.GZ43_512545	AF111855	Agrobacterium tumefaciens RNA polymerase alpha subunit (rpoA) gene, complete cds	2.00E-06
685	3596.P07.GZ43_512593	L40817	Homo sapiens muscle-specific DNase I-like (DNLI1) gene, exons 1-9, complete cds	3.00E-06
686	3596.P08.GZ43_512609	M14505	Human (clone PSK-13) cyclin-dependent protein kinase mRNA, complete cds,	5.00E-06
687	3596.P10.GZ43_512641	M73770	P.falciparum RNA polymerase III largest subunit gene, complete cds	2.90E-05

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
688	3596.P21.GZ43_512817	S82725	NPM/ALK-fusion gene (translocation breakpoint) [human, lymphoma cells SU-DHL-1, Genomic, 1679 nt]	1.00E-07
689	3599.A04.GZ43_512914	X83212	H.sapiens tryptophan hydroxylase gene, promoter region	5.50E-07
690	3599.A23.GZ43_513218	U05259	Human MB-1 gene, complete cds	2.10E-05
691	3599.B15.GZ43_513091	AF277068	HIV-1 clone QH0791 from Trinidad and Tobago, envelope protein (env) gene, complete cds	6.10E-07
692	3599.B16.GZ43_513107	M60517	Chicken vitronectin receptor alpha subunit mRNA, complete cds	4.00E-06
693	3599.C03.GZ43_512900	AB021267	Arabidopsis thaliana copia-like retrotransposon AtRE2-2 gene for polyprotein, complete cds	2.00E-06
694	3599.C17.GZ43_513124	U28055	Homo sapiens hepatocyte growth factor-like protein homolog mRNA, partial cds	3.00E-06
695	3599.D03.GZ43_512901	L43550	Buchnera aphidicola anthranilate synthase small subunit (trpG) gene, anthranilate synthase large subunit (trpE) gene, complete cds	3.00E-06
696	3599.D05.GZ43_512933	AL023779	S.pombe chromosome II cosmid c244	2.00E-06
697	3599.D07.GZ43_512965	AL391223	Human chromosome 14 DNA sequence Partial sequence from BAC R-325N7_PCR1 of library RPCT-11 from chromosome 14 of Homo sapiens (Human), complete sequence	5.00E-06
698	3599.D10.GZ43_513013	AF064079	Plasmodium gallinaceum endochitinase precursor, mRNA, complete cds	1.70E-07
699	3599.E01.GZ43_512870	U09184	Buchnera aphidicola ferredoxin-NADP reductase (fpr1) gene, partial cds; anthranilate synthase large subunit (trpE) and anthranilate synthase small subunit (trpG) genes, complete cds; heat shock protein (hslU) gene, partial cds; and unknown gene	9.60E-07
700	3599.E05.GZ43_512934	X60145	Human J-alpha segment J-alpha FR9 mRNA for J-alpha region of T-cell receptor	1.20E-05
701	3599.F17.GZ43_513127	U27037	Fistulina hepatica mitochondrial small subunit ribosomal RNA, mitochondrial gene, partial sequence	2.00E-06
702	3599.F24.GZ43_513239	Z78414	Caenorhabditis elegans cosmid W09D12, complete sequence	5.00E-06
703	3599.H05.GZ43_512937	AF032891	Camponotus consobrinus microsatellite-containing sequence Ccon12	2.10E-08
704	3599.H23.GZ43_513225	AB024553	Bacillus halodurans DNA, complete and partial cds, strain:C-125	4.70E-07
705	3599.J11.GZ43_513035	AB025112	Xenopus laevis XGC-2 mRNA for guanylyl cyclase-2, complete cds	3.00E-06
706	3599.K02.GZ43_512892	AJ224474	Borrelia burgdorferi left chromosomal subtelomeric region (truA gene)	3.00E-06

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
707	3599.K04.GZ43_512924	X99710	L.lactis ORF, genes homologous to vsf-1 and pepF2 and gene encoding protein homologous to methyltransferase	5.00E-06
708	3599.K23.GZ43_513228	AF074247	Homo sapiens neuronal delayed-rectifier voltage-gated potassium channel splice variant (KCNQ2) mRNA, complete cds	8.00E-07
709	3599.L04.GZ43_512925	X59773	Pisum sativum mRNA for P protein, a part of glycine cleavage complex	1.40E-05
710	3599.L15.GZ43_513101	U34282	Rattus norvegicus fast skeletal muscle sarcoplasmic reticulum Ca-ATPase (SERCA1) gene, 5'-flanking sequence	2.00E-06
711	3599.M04.GZ43_512926	AK018953	Mus musculus adult male testis cDNA, RIKEN full-length enriched library, clone:1700111D04, full insert sequence	2.30E-11
712	3599.M22.GZ43_513214	AB052179	Macaca fascicularis brain cDNA, clone:QnpA-21934	4.70E-07
713	3599.M24.GZ43_513246	AE003394	Drosophila melanogaster genomic scaffold 142000013386028, complete sequence	7.30E-07
714	3599.N09.GZ43_513007	X16362	Rat SPI-2 serine protease inhibitor gene	1.19E-04
715	3599.N16.GZ43_513119	X92421	X.laevis mRNA for RNA helicase p54	3.00E-06
716	3599.N20.GZ43_513183	M59447	Drosophila melanogaster Sex-lethal (Sxl) mRNA, complete cds	2.00E-06
717	3599.N24.GZ43_513247	AC005485	Homo sapiens PAC clone RP5-998M2 from 7q33-q35, complete sequence	2.00E-07
718	3599.O06.GZ43_512960	AJ131667	Escherichia coli plasmid pSFO157	2.00E-06
719	3599.O17.GZ43_513136	X96607	M.musculus IgH 3' alpha enhancer DNA	8.10E-05
720	3599.P05.GZ43_512945	X77111	N.tabacum chi-V gene	1.50E-07
721	3602.A09.GZ43_513378	AF015303	Xenopus laevis small GTPase Ran binding protein 1 mRNA, complete cds	1.10E-05
722	3602.B18.GZ43_513523	L18892	Tetrahymena thermophila histone (H2A.1) gene, complete cds	5.70E-07
723	3602.B21.GZ43_513571	BC005233	Homo sapiens, clone MGC:12257 IMAGE:3950129, mRNA, complete cds	1.60E-10
724	3602.B22.GZ43_513587	X71765	P. falciparum gene for Ca2+ - ATPase	1.00E-06
725	3602.C24.GZ43_513620	AL080106	Homo sapiens mRNA; cDNA DKFZp566O053 (from clone DKFZp566O053)	2.00E-06
726	3602.D06.GZ43_513333	AF098970	Phascolus vulgaris NBS-LRR-like protein cD7 (CO-2) mRNA, partial cds	1.70E-07
727	3602.D11.GZ43_513413	M59770	P.falciparum calmodulin gene, complete cds	2.20E-07
728	3602.E04.GZ43_513302	X53582	Zea mays ZMPMS1 gene for 19 kDa zein protein	1.30E-05
729	3602.E06.GZ43_513334	L38718	Providencia stuartii (clone pSK.aarP) transcriptional activator (aarP) gene, complete cds	7.90E-07
730	3602.E13.GZ43_513446	U58106	Blomia tropicalis allergen mRNA, complete cds	1.70E-07

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
731	3602.E21.GZ43_513574	M15085	T.brucei expressed copy of the ILTat 1.3 variable surface glycoprotein gene, 5' flank	2.90E-07
732	3602.F12.GZ43_513431	X64802	H.sapiens F8 mRNA for Interleukin-1-like species	3.40E-58
733	3602.G03.GZ43_513288	AF036148	Danio rerio NeuroD (nrd) mRNA, complete cds	2.00E-06
734	3602.G17.GZ43_513512	U41106	Caenorhabditis elegans cosmid W06A11	1.30E-05
735	3602.I07.GZ43_513354	AF000941	Mus musculus DNase I hypersensitive sites 2-6 of locus control region (LCR) for T-cell receptor alpha chain (TCRa) gene	1.20E-05
736	3602.I11.GZ43_513418	AL133620	Homo sapiens mRNA; cDNA DKFZp434F0621 (from clone DKFZp434F0621)	3.00E-06
737	3602.I15.GZ43_513482	U23479	Dictyostelium discoideum phosphatidylinositol 4-kinase (PIK4) mRNA, complete cds	8.00E-07
738	3602.J13.GZ43_513451	AK025319	Homo sapiens cDNA: FLJ21666 fis, clone COL08915	3.30E-07
739	3602.K03.GZ43_513292	X85811	S.cerevisiae tRNA-Leu, and ORF's N2212, N2215, N2219, N2223, N2227, N2231	1.10E-05
740	3602.K06.GZ43_513340	AF133052	Walleye epidermal hyperplasia virus type 2 long terminal repeat, complete sequence; gag polyprotein (gag-pol) gene, complete cds; pol polyprotein (gag-pol) gene, partial cds; envelope polyprotein (env) and cyclin D homolog genes, complete cds; and unkn>	4.00E-06
741	3602.L20.GZ43_513565	M62717	Human CSP-B gene flanking sequence	1.10E-05
742	3602.N03.GZ43_513295	Z81126	Caenorhabditis elegans cosmid T22E6, complete sequence	5.70E-05
743	3602.N06.GZ43_513343	U62503	Human OBR gene, intron sequence immediately adjacent to the 5' end of coding exon 17	1.00E-06
744	3605.A15.gz43_513858	Z46507	Bovine herpesvirus type 4 genomic DNA region (V.TEST)	5.00E-06
745	3605.C16.gz43_513876	AF282517	Homo sapiens clone 10ptel_c617 sequence	9.40E-08
746	3605.E19.gz43_513926	Z22923	M.musculus alpha2 (IX) collagen gene, complete CDS	2.10E-05
747	3605.G13.gz43_513832	AJ132752	Gadus morhua mRNA for beta2-microglobulin, clone b3	1.30E-05
748	3605.H10.gz43_513785	AF257480	Rana temporaria microsatellite SB80 sequence	4.10E-09
749	3605.H21.gz43_513961	X63507	M.musculus HOX-3.5 gene	7.80E-05
750	3605.I19.gz43_513930	AK002100	Homo sapiens cDNA FLJ11238 fis, clone PLACE1008532	3.30E-11
751	3605.J16.gz43_513883	AF039197	Gallus gallus Pax-9 gene, putative 5' regulatory sequence	1.00E-07

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
752	3605.K19.gz43_513932	X63853	<i>S.cerevisiae</i> MAT locus genes BUD5, mat-alpha1, mat-alpha2, YCR724 and YCR725	8.00E-06
753	3605.M17.gz43_513902	M30931	Simian immunodeficiency virus (SIV) proviral, complete genome	3.70E-05
754	3605.N04.gz43_513695	AF169388	<i>Mus musculus</i> alpha 4 collagen IV (Col4a4) mRNA, complete cds	8.90E-05
755	3605.N09.gz43_513775	AF029111	<i>Adelius</i> sp. 16S ribosomal RNA gene, mitochondrial gene for mitochondrial RNA, partial sequence	2.80E-07
756	3605.N12.gz43_513823	BC000358	Homo sapiens, protein kinase, AMP-activated, gamma 1 non-catalytic subunit, clone MGC:8666 IMAGE:2964434, mRNA, complete cds	3.90E-47
757	3605.N16.gz43_513887	X95301	<i>D. rerio</i> mRNA for HER-5 protein	1.00E-06
758	3608.B06.gz43_514099	X00004	<i>t.aurus</i> gene encoding pituitary glycoprotein hormone alpha subunit, exons 3 & 4	6.30E-08
759	3608.B12.gz43_514195	X00525	Mouse 28S ribosomal RNA	3.10E-13
760	3608.B24.gz43_514387	AF269848	<i>Staphylococcus epidermidis</i> strain SR1 clone step.102606 genomic sequence	2.00E-06
761	3608.C18.gz43_514292	BC000387	Homo sapiens, U6 snRNA-associated Sm-like protein, clone MGC:8433 IMAGE:2821171, mRNA, complete cds	2.50E-10
762	3608.E17.gz43_514278	BC008245	Homo sapiens, clone IMAGE:3875012, mRNA	1.00E-06
763	3608.E20.gz43_514326	U86646	<i>Ailurus fulgens</i> beta casein gene, exon 7, partial cds	4.70E-07
764	3608.F13.gz43_514215	AF125672	Homo sapiens silencing mediator of retinoic acid and thyroid hormone receptor extended isoform (SMRTE) mRNA, complete cds	2.00E-06
765	3608.G09.gz43_514152	AE001066	<i>Archaeoglobus fulgidus</i> section 41 of 172 of the complete genome	4.00E-06
766	3608.H05.gz43_514089	AJ224981	<i>Mus musculus</i> calpain 3 gene, exon 1	3.00E-06
767	3608.H14.gz43_514233	AE007394	<i>Streptococcus pneumoniae</i> section 77 of 194 of the complete genome	3.20E-05
768	3608.H18.gz43_514297	Z36046	<i>S.cerevisiae</i> chromosome II reading frame ORF YBR177c	7.00E-06
769	3608.J17.gz43_514283	AF024648	<i>Arabidopsis thaliana</i> receptor-like serine/threonine kinase (RKF1) mRNA, complete cds	8.00E-06
770	3608.J24.gz43_514395	AJ002258	<i>Rattus Norvegicus</i> mRNA for Prx3A protein	3.60E-07
771	3608.K03.gz43_514060	M83199	<i>Simmondsia chinensis</i> stearyl-acyl carrier protein desaturase mRNA, complete cds	2.50E-07
772	3608.K14.gz43_514236	AK026999	Homo sapiens cDNA: FLJ23346 fis, clone HEP13716	2.00E-06
773	3608.L07.gz43_514125	M32684	Homo sapiens ITGB3 gene, intron 13, fragment B, partial sequence	3.60E-07



Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
774	3608.L14.gz43_514237	Z34845	H.sapiens serotonin transporter gene	8.60E-07
775	3608.N09.gz43_514159	AK022341	Homo sapiens cDNA FLJ12279 fis, clone MAMMA1001743, weakly similar to Y BOX BINDING PROTEIN-1	2.00E-06
776	3608.N19.gz43_514319	M15085	T.brucei expressed copy of the ILT'at 1.3 variable surface glycoprotein gene, 5' flank	7.80E-08
777	3608.N20.gz43_514335	AF026169	Homo sapiens SALF (SALF) mRNA, complete cds	1.00E-05
778	3608.O04.gz43_514080	U85193	Human nuclear factor I-B2 (NFIB2) mRNA, complete cds	7.10E-07
779	3608.P22.gz43_514369	AF124241	Callerya australis chloroplast tRNA-Leu (trnL) gene, intron sequence	3.90E-07
780	3611.A17.gz43_514658	X01412	Drosophila melanogaster genes for tRNA-Val and tRNA-Pro (90BC tRNA locus)	2.00E-06
781	3611.B11.gz43_514563	AL049938	Homo sapiens mRNA; cDNA DKFZp564P1916 (from clone DKFZp564P1916); partial cds	9.80E-10
782	3611.B16.gz43_514643	M86514	Rat proline-rich protein mRNA, 3' end	1.30E-05
783	3611.C09.gz43_514532	U55950	Pleurodeles waltl cytochrome b (CYT-b) gene, mitochondrial gene encoding mitochondrial protein, partial cds	2.00E-06
784	3611.E07.gz43_514502	AF261009	Lethrinus miniatus clone 89rte, microsatellite sequence	1.70E-12
785	3611.E12.gz43_514582	M60200	Rat vitamin D binding protein gene, exons 5 and 6	1.50E-05
786	3611.E20.gz43_514710	BC002458	Homo sapiens, clone IMAGE:3343171, mRNA, partial cds	2.00E-06
787	3611.F15.gz43_514631	U28328	Bos taurus dinucleotide repeat RM154, tandem repeat region	4.30E-27
788	3611.H10.gz43_514553	AE003147	Drosophila melanogaster genomic scaffold 142000013385388, complete sequence	6.00E-07
789	3611.H22.gz43_514745	X16135	Human mRNA for novel heterogeneous nuclear RNP protein, L protein	7.00E-06
790	3611.I04.gz43_514458	AK001460	Homo sapiens cDNA FLJ10598 fis, clone NT2RP2004841	5.10E-44
791	3611.I13.gz43_514602	M58380	Arabidopsis thaliana peroxidase (neutral, prxCa) gene, complete cds	3.00E-06
792	3611.J04.gz43_514459	S81486	p53 [alternatively spliced, intron 9] [human, Genomic Mutant, 133 nt]	1.20E-07
793	3611.J15.gz43_514635	AC008240	Leishmania major chromosome 22 clone L9259 strain Friedlin, complete sequence	4.90E-05
794	3611.J17.gz43_514667	Z17425	Lilium speciosum for two putative cds's	8.90E-07
795	3611.J22.gz43_514747	U60736	Human IgHC locus intergenic sequence	4.60E-07
796	3611.K01.gz43_514412	AE001377	Plasmodium falciparum chromosome 2, section 14 of 73 of the complete sequence	3.00E-06
797	3611.K12.gz43_514588	X02367	Glaucoma chattoni rDNA 3' NTS	9.80E-08
798	3611.L22.gz43_514749	U19361	Petromyzon marinus neurofilament subunit NF-180 mRNA, complete cds	5.40E-08

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
799	3611.M18.gz43_514686	X95301	D.erio mRNA for HER-5 protein	1.00E-06
800	3611.M24.gz43_514782	AF010239	Caenorhabditis elegans glutathione S-transferase (CeGST1) mRNA, complete cds	7.70E-07
801	3611.N01.gz43_514415	L19300	Staphylococcus aureus DNA sequence encoding three ORFs, complete cds; prophage phi-11 sequence homology, 5' flank	1.00E-06
802	3611.N09.gz43_514543	U50382	Danio rerio beta and alpha globin genes, partial cds	7.00E-06
803	3611.O16.gz43_514656	AB056785	Macaca fascicularis brain cDNA clone:QmpA-11655, full insert sequence	6.60E-07
804	3611.P08.gz43_514529	AK026905	Homo sapiens cDNA: FLJ23252 fis, clone COL04668	8.00E-06
805	3614.C18.gz43_515060	AF239178	Paracoccidioides brasiliensis lon proteinase gene, complete cds; nuclear gene for mitochondrial product	5.00E-06
806	3614.D14.gz43_514997	AB017511	Hydra magnipapillata mRNA for PLC-betaH1, complete cds	1.20E-05
807	3614.D21.gz43_515109	L10713	Pig trinucleotide repeat	1.80E-05
808	3614.E06.gz43_514870	X99739	M.musculus mRNA for UBC9 protein, containing ubiquitin box	9.10E-07
809	3614.F22.gz43_515127	AK021490	Homo sapiens cDNA FLJ11428 fis, clone HEMBA1001071, highly similar to PROCOLLAGEN ALPHA 1(III) CHAIN PRECURSOR	2.00E-06
810	3614.G20.gz43_515096	M86514	Rat proline-rich protein mRNA, 3' end	1.30E-05
811	3614.H09.gz43_514921	AF068289	Homo sapiens HDCMD34P mRNA, complete cds	6.60E-11
812	3614.H22.gz43_515129	X62423	P.falciparum pol delta gene for DNA polymerase delta	4.00E-06
813	3614.J07.gz43_514891	X81027	H.sapiens tal-1 DNA	1.30E-05
814	3614.K22.gz43_515132	X63073	Pseudanabaena sp. cpeBA operon encoding phycoerythrin beta and alpha subunits	1.60E-05
815	3614.L13.gz43_514989	V01561	Mouse dispersed repetitive DNA sequences of the R-family and simple sequence DNA; member of the B1 family of mouse dispersed repetitive DNA sequences	3.00E-06
816	3614.M08.gz43_514910	AF272983	Homo sapiens SRC tyrosine kinase gene, exons 1alpha and 1a, alternatively spliced	4.00E-06
817	3614.O02.gz43_514816	X58913	Mitochondrion Drosophila cucurculis ND2 and COI genes (partial) and genes for tRNA-Trp, tRNA-Tyr, and tRNA-Cys	8.50E-08
818	3614.O07.gz43_514896	AL031538	S.pombe chromosome III cosmid c1906	9.80E-07
819	3614.O16.gz43_515040	AB056785	Macaca fascicularis brain cDNA clone:QmpA-11655, full insert sequence	2.00E-06
820	3614.P11.gz43_514961	X91656	M.musculus Srp20 gene	4.60E-05

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
821	3614.P16.gz43_515041	Z58907	H.sapiens CpG island DNA genomic MseI fragment, clone 116a6, forward read cpg116a6.fl1a	3.20E-70
822	3617.B16.gz43_515411	AF098275	Homo sapiens PS12TOM20 pseudogene, complete sequence	1.10E-67
823	3617.C21.gz43_515492	AJ009913	Bos taurus plp gene	3.40E-05
824	3617.F10.gz43_515319	L07487	Bradyrhizobium japonicum heme-copper oxidase subunit I homolog (fixN), cytochrome c (fixO), transmembrane proteins (fixO and fixQ) di-heme cytochrome c (fixP) and fixG genes, complete cds	6.70E-05
825	3617.H16.gz43_515417	X54192	O.sativa GlnB-2 gene for glutelin	2.00E-06
826	3617.I01.gz43_515178	AL513316	Human DNA sequence from clone RP11-522O3 on chromosome 10, complete sequence [Homo sapiens]	7.20E-08
827	3617.L16.gz43_515421	AE007662	Clostridium acetobutylicum ATCC824 section 150 of 356 of the complete genome	3.00E-06
828	3617.L21.gz43_515501	AL031538	S.pombe chromosome III cosmid c1906	1.00E-06
829	3617.M08.gz43_515294	X64802	H.sapiens F8 mRNA for Interleukin-1-like species	3.40E-58
830	3617.M13.gz43_515374	Z79239	H.sapiens flow-sorted chromosome 6 TaqI fragment, SC6pA26F6	1.10E-07
831	3617.N05.gz43_515247	AF387666	Mandrillus cytomegalovirus strain OCOM6-2 glycoprotein B (gB) gene, partial cds	1.00E-06
832	3617.N10.gz43_515327	AB017511	Hydra magnipapillata mRNA for PLC-betaH1, complete cds	1.10E-05
833	3617.N14.gz43_515391	AJ249346	Mus musculus Ankrd2 gene for ankyrin repeat domain 2 (stretch responsive muscle), exons 1-9	1.00E-05
834	3617.N19.gz43_515471	U27037	Fistulina hepatica mitochondrial small subunit ribosomal RNA, mitochondrial gene, partial sequence	2.00E-06
835	3617.P11.gz43_515345	AK002100	Homo sapiens cDNA FLJ11238 fis, clone PLACE1008532	1.20E-13
836	3617.P12.gz43_515361	U04860	Rattus norvegicus Sprague-Dawley Ah receptor mRNA, complete cds	8.00E-05
837	3617.P13.gz43_515377	AE007356	Streptococcus pneumoniae section 39 of 194 of the complete genome	3.80E-05
838	3620.B03.gz43_515810	AF238884	Botrytis virus F, complete genome	6.00E-06
839	3620.B24.gz43_516146	AF244812	Homo sapiens SCAN domain-containing protein 2 (SCAND2) gene, complete cds, alternatively spliced	1.30E-07
840	3620.E12.gz43_515957	X95301	D.rerio mRNA for HER-5 protein	1.00E-06
841	3620.E13.gz43_515973	X52289	Human (D21S167) DNA segment containing (GT)19 repeat	2.50E-19
842	3620.E17.gz43_516037	AJ002414	Arabidopsis thaliana mRNA for a hnRNP-like protein	9.70E-08

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
843	3620.E19.gz43_516069	X16982	Drosophila melanogaster micropia-Dm11 3'flanking DNA	2.70E-07
844	3620.E23.gz43_516133	Z49438	S. cerevisiae chromosome X reading frame ORF YJL163c	3.00E-06
845	3620.E24.gz43_516149	M75883	Human sterol carrier protein X/sterol carrier protein 2 mRNA, complete cds	8.00E-06
846	3620.G17.gz43_516039	U92971	Human protease-activated receptor 3 (PAR3) mRNA, complete cds	3.80E-07
847	3620.G23.gz43_516135	X66979	X. laevis mRNA XLFLI	1.60E-05
848	3620.J18.gz43_516058	U37373	Xenopus laevis tail-specific thyroid hormone up-regulated (gene 5) mRNA, complete cds	3.00E-06
849	3620.K19.gz43_516075	U31780	Human papillomavirus type 22, complete genome	5.00E-06
850	3620.K24.gz43_516155	M95627	Homo sapiens angio-associated migratory cell protein (AAMP) mRNA, complete cds	6.00E-06
851	3620.O23.gz43_516143	L11172	Plasmodium falciparum RNA polymerase I gene, complete cds	1.00E-05
852	3623.B07.gz43_516258	AF132745	Mus musculus Sox2 gene, regulatory region sequence	7.70E-07
853	3623.B03.gz43_516197	X82566	M.musculus glyT1 gene (exon 0a)	1.80E-09
854	3623.E15.gz43_516389	AF104420	Porcine transmissible gastroenteritis virus RNA dependent RNA polymerase gene, partial cds; virus envelope protein spike (S), envelope protein (sm), envelope protein (M), and nucleoprotein (N) genes, complete cds; and unknown genes	2.90E-05
855	3623.F03.gz43_516198	AJ009936	Homo sapiens mRNA for nuclear hormone receptor PRR1	1.70E-05
856	3623.F20.gz43_516470	U22657	Mus musculus genomic locus related to cellular morphology	5.80E-05
857	3623.G14.gz43_516375	AB035309	Paramecium caudatum PcTERT mRNA for telomerase reverse transcriptase, complete cds	3.00E-06
858	3623.H07.gz43_516264	Z17324	Homo sapiens of MUC1 gene encoding Mucin	1.80E-07
859	3623.H10.gz43_516312	AB033070	Homo sapiens mRNA for KIAA1244 protein, partial cds	2.80E-05
860	3623.H23.gz43_516520	AF131763	Homo sapiens clone 25232 mRNA sequence	1.70E-05
861	3623.I08.gz43_516281	M60421	Human cytochrome P450sc gene, 5' end and promoter region	2.80E-05
862	3623.I11.gz43_516329	AK013191	Mus musculus 10, 11 days embryo cDNA, RIKEN full-length enriched library, clone:2810429I04, full insert sequence	3.00E-06
863	3623.L05.gz43_516236	AJ131991	Linum usitatissimum target sequence for LIS-1 insertion in PI	3.00E-06

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
864	3623.L24.gz43_516540	U09377	Arabidopsis thaliana GF14chi isoform (GRF1) gene, complete cds	3.00E-06
865	3623.M10.gz43_516317	AF071743	Homo sapiens topoisomerase II alpha (TOP2A) gene, exons 25, 26, and 27	4.00E-06
866	3623.N23.gz43_516526	U57489	Bubacterium sp. VPI 12708 bile acid-inducible operon bile acid-coenzyme A ligase (baiB), BaiC, BaiD, bile acid 7-alpha dehydratase (baiE), 3-alpha hydroxysteroid dehydrogenase (baiA2), BaiF, bile acid transporter (baiG), NADH:flavin oxidoreductase (bai>	3.70E-05
867	3623.P22.gz43_516512	U37761	Human H1 histamine receptor gene, 5'-flanking region	1.40E-12
868	3626.A10.gz43_516689	D30745	Xenopus laevis MRP RNA gene	2.00E-07
869	3626.C16.gz43_516787	AF241271	Bos taurus ZFY gene, intron	1.60E-08
870	3626.E07.gz43_516645	AF053496	Caenorhabditis elegans beta chain spectrin homolog Smal1 (smal1) mRNA, complete cds	2.00E-06
871	3626.F03.gz43_516582	AJ009771	Homo sapiens mRNA for putative RING finger protein, partial	2.00E-06
872	3626.G01.gz43_516551	BC010926	Homo sapiens, Similar to H4 histone family, member A, clone MGC:13512 IMAGE:4273904, mRNA, complete cds	1.00E-43
873	3626.I20.gz43_516857	AK025762	Homo sapiens cDNA: FLJ22109 fis, clone HEP18091 (156)=G surface antigen {3' region, restriction fragment EG4} [Parametium primaurelia, Genomic, 407 nt]	5.80E-07
874	3626.I23.gz43_516905	S55615	Plasmodium falciparum chromosome 2, section 35 of 73 of the complete sequence	4.00E-06
875	3626.M13.gz43_516749	AE001398	Homo sapiens clone HQ0452 PRO0452 mRNA, partial cds	3.10E-07
876	3626.M15.gz43_516781	AF090925	H.sapiens CpG island DNA genomic MseI fragment, clone 116a6, forward read	2.90E-70
877	3626.N07.gz43_516654	Z58907	Rattus norvegicus clathrin assembly protein short form (CALM) mRNA, complete cds	8.90E-08
878	3626.N24.gz43_516926	AF041373	Mouse mRNA for protein C, complete cds	5.00E-06
879	3626.O08.gz43_516671	D10445	Homo sapiens (subclone 6_h1 from P1 H21) DNA sequence	2.20E-07
880	3626.P11.gz43_516720	L48479	Chicken hsp90 gene for 90 kDa-heat shock protein 5'-end	3.80E-05
881	3626.P14.gz43_516768	X15028	Mus musculus pre-T cell receptor alpha-type chain precursor mRNA, complete cds	4.00E-06
882	3629.A16.gz43_517169	U16958	Drosophila melanogaster micropia-Dm11 3'flanking DNA	2.50E-07
883	3629.B14.gz43_517138	X16982		

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
884	3629.C14.gz43_517139	Z22537	C.parvum precursor of oocyst wall protein	5.00E-06
885	3629.E01.gz43_516933	D00621	Sus scrofa gene for follicle stimulation hormone beta subunit, exons 1, 2, 3, complete cds	3.50E-05
886	3629.E20.gz43_517237	AH3006900	Sulfolobus solfataricus section 259 of 272 of the complete genome	9.00E-06
887	3629.F24.gz43_517302	Y10531	Clostridium perfringens sod gene for superoxide dismutase	2.00E-06
888	3629.H10.gz43_517080	J03654	Human immunodeficiency virus type 2, isolate HIV2FG	8.00E-06
889	3629.H12.gz43_517112	AF017266	Danio rerio glutamate decarboxylase (GAD67) mRNA, partial cds	6.50E-07
890	3629.I11.gz43_517097	AF020810	Salmonella enterica VirK (virK), Mig-14 (mig-14), NxiA (nxiA), TctE (tctE), TctD (tctD), TctC (tctC), TctB (tctB), and TctA (tctA) genes, complete cds, and O360 (o360) gene, partial cds	3.00E-06
891	3629.II6.gz43_517177	AF007643	Clostridium acetobutylicum ATCC824 section 131 of 356 of the complete genome	4.40E-05
892	3629.J03.gz43_516970	AB017511	Hydra magnipapillata mRNA for PLC-betaH1, complete cds	1.10E-05
893	3629.J07.gz43_517034	M20782	Human alpha-2-plasmin inhibitor gene, exons 2 to 5	2.90E-11
894	3632.C11.gz43_517475	AF026148	Perilla frutescens beta-ketoacyl-ACP synthase I (KAS I) mRNA, complete cds	1.00E-06
895	3632.C17.gz43_517571	U50534	Human BRCA2 region, mRNA sequence CG003	1.00E-05
896	3632.F07.gz43_517414	M12036	Human tyrosine kinase-type receptor (HER2) gene, partial cds	4.70E-10
897	3632.G01.gz43_517319	AC006621	Caenorhabditis elegans cosmid CS2A10, complete sequence	3.40E-05
898	3632.I20.gz43_517625	AK024381	Homo sapiens cDNA FLJ14319 fis, clone PLACE3000406	9.00E-06
899	3632.K20.gz43_517627	M27634	Vaccinia virus P4a major core protein gene, complete cds	9.60E-05
900	3632.M08.gz43_517437	X75304	H.sapiens gintonin mRNA	8.00E-06
901	3632.M13.gz43_517517	U18191	Human HLA class I genomic survey sequence	3.20E-07
902	3632.M19.gz43_517613	AF012131	Homo sapiens brachyury variant B (TBX1) mRNA, complete cds	3.70E-07
903	3632.N13.gz43_517518	AF287491	Oncorhynchus mykiss MHC class I heavy chain precursor (Onmy-UBA) mRNA, Onmy-UBA*601 allele, complete cds	2.00E-06
904	3632.N21.gz43_517646	X62423	P.falciparum pol delta gene for DNA polymerase delta	4.00E-06
905	3632.O06.gz43_517407	BC009868	Homo sapiens, replication protein A3 (14kD), clone MGC:16404 IMAGE:3940438, mRNA, complete cds	1.40E-18

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Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
906	3632.P07.gz43_517424	AF001066	Archaeoglobus fulgidus section 41 of 172 of the complete genome	3.00E-06
907	3635.A06.gz43_517777	AK005546	Mus musculus adult female placenta cDNA, RIKEN full-length enriched library, clone:1600027G01; full insert sequence	1.40E-07
908	3635.A08.gz43_517809	Z49280	S.cerevisiae chromosome X reading frame ORF YJL005w	6.00E-06
909	3635.A13.gz43_517889	AF143236	Homo sapiens apoptosis related protein APR 2 mRNA, complete cds	2.00E-06
910	3635.D07.gz43_517796	M58150	Bovine lactoperoxidase (LPO) mRNA, complete cds	3.10E-05
911	3635.F01.gz43_517702	Y19128	Homo sapiens enteropeptidase gene, exon 6	3.00E-09
912	3635.F06.gz43_517782	X63073	Pseudanabaena sp. cpeBA operon encoding phycoerythrin beta and alpha subunits	1.50E-05
913	3635.F10.gz43_517846	AF107688	Aedes aegypti clone 431 Fellal family of SINES	3.50E-05
914	3635.H20.gz43_518008	AE000613	Helicobacter pylori 26695 section 91 of 134 of the complete genome	1.10E-05
915	3635.J06.gz43_517786	U15018	Dugbe virus L protein gene, complete cds	1.10E-05
916	3635.J09.gz43_517834	X85444	G.pallida repetitive DNA element	2.10E-08
917	3635.K05.gz43_517771	AF090432	Danio rerio serrateB mRNA, complete cds	4.00E-06
918	3635.K06.gz43_517787	AJ276631	Capsicum annuum partial kn gene for Knolle protein, promoter region	6.10E-07
919	3635.M18.gz43_517981	AL591498	Human DNA sequence from clone RP11-113L12 on chromosome 13, complete sequence [Homo sapiens]	1.40E-05
920	3635.O01.gz43_517711	AF081788	Homo sapiens putative spliceosome associated protein mRNA, complete cds	3.70E-30
921	3635.O14.gz43_517919	X72224	S.cerevisiae genes HSS1, NPL4 and HSP	6.00E-06
922	3635.P17.gz43_517968	AF242307	Euphorbia esula sucrose transport protein mRNA, complete cds	2.90E-10
923	3635.P18.gz43_517984	AF078780	Caenorhabditis elegans cosmid C04F2, complete sequence	1.74E-04
924	3638.A02.gz43_518097	MI7988	Spiroplasma virus 4 (SpV4) replicative form, complete genome	4.00E-06
925	3638.A24.gz43_518449	AF064079	Plasmodium gallinaceum endochitinase precursor, mRNA, complete cds	1.60E-07
926	3638.F15.gz43_518310	AJ297538	Homo sapiens partial RARA gene, intron 2	4.00E-06
927	3638.H07.gz43_518184	AK026258	Homo sapiens cDNA: FLJ22605 fig, clone HSI04743	2.00E-06
928	3638.J09.gz43_518218	U89651	Homo sapiens matrix metalloproteinase MMP Rasi-1 gene, promoter region	8.10E-08
929	3638.K06.gz43_518171	AL139329	Human DNA sequence from clone RP11-228P1 on chromosome 6, complete sequence [Homo sapiens]	4.40E-11

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
930	3638.L10.gz43_518236	D26532	Mouse mRNA for transcription factor PEBP2aB2, complete cds	2.00E-08
931	3638.N05.gz43_518158	X62294	B.taurus mRNA for adrenal angiotensin II type-1 receptor	9.00E-06
932	3643.D21.gz43_518788	U17010	Allomyces macrogynus mitochondrion NADH dehydrogenase subunit 5 (nad5) gene, complete cds	1.80E-05
933	3643.E24.gz43_518837	AL022342	Human DNA sequence from clone RP1-29M10 on chromosome 20, complete sequence [Homo sapiens]	6.70E-05
934	3643.F07.gz43_518566	M73962	Bovine pregnancy-associated glycoprotein 1 mRNA, complete cds	6.00E-06
935	3643.G20.gz43_518775	AF191214	Homo sapiens isovaleryl dehydrogenase (IVD) gene, exons 1-3	1.00E-05
936	3643.G24.gz43_518839	AK025682	Homo sapiens cDNA: FLJ22029 fis, clone HEP08661	6.00E-06
937	3643.H09.gz43_518600	AK024381	Homo sapiens cDNA FLJ14319 fis, clone PLACE3000406	1.70E-05
938	3643.I01.gz43_518473	AF000306	Brassica napus steroid sulfotransferase 2 gene, complete cds	3.00E-06
939	3643.I02.gz43_518489	X58433	B.subtillis cad gene for lysine decarboxylase	2.30E-05
940	3643.I18.gz43_518745	M14872	Mouse GnRH-GAP gene encoding gonadotropin-releasing hormone and GnRH-associated peptide (GAP)	4.00E-06
941	3643.I24.gz43_518841	BC003813	Mus musculus, clone MGC:6139	
942	3643.K06.gz43_518555	AL050124	IMAGE:3487295, mRNA, complete cds	2.30E-07
943	3643.L01.gz43_518476	AJ278429	Homo sapiens mRNA; cDNA DKFZp586E151 (from clone DKFZp586E151)	1.60E-07
944	3643.N24.gz43_518846	BC006511	Mus musculus partial Prkar1a gene for cAMP-dependent protein kinase regulatory subunit R1alpha, exons 8-10 and 3'UTR	3.00E-06
945	3643.O16.gz43_518719	AE002303	Homo sapiens, clone IMAGE:3010441, mRNA	1.00E-05
946	3643.O18.gz43_518751	V00248	Chlamydia muridarum, section 34 of 85 of the complete genome	1.10E-05
947	3643.O21.gz43_518799	AE000614	Drosophila gene for yolk protein I (vitellogenin)	2.00E-06
948	3643.P13.gz43_518672	Y17693	Helicobacter pylori 26695 section 92 of 134 of the complete genome	1.40E-05
949	3643.P14.gz43_518688	AF109352	Bungarus multicinctus gene encoding alpha-bungarotoxin, V31 variant	2.00E-07
950	3646.A07.gz43_518945	X55137	Euperipatoides rowelli microsatellite P18 sequence	8.80E-10
951	3646.A09.gz43_518977	AF074963	H. giganteus type II restriction-modification system HgiBI	3.00E-06
			Rattus norvegicus endothelin-B receptor (EDNRB) gene, partial cds	2.10E-07



Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
952	3646.A12.gz43_519025	AF176208	Homo sapiens EcoRI-HindIII fragment upstream of exon 1 of the c-myc gene	1.60E-05
953	3646.A13.gz43_519041	X89445	O. chalybea DNA for narB gene and partial ORFs	4.00E-05
954	3646.B20.gz43_519154	M86514	Rat proline-rich protein mRNA, 3' end	1.60E-05
955	3646.C06.gz43_518931	Z71180	Caenorhabditis elegans cosmid F22E12, complete sequence	2.03E-04
956	3646.C16.gz43_519091	U73608	Hepatitis B virus, genome 7648 with G->A hypermutations	2.30E-05
957	3646.E02.gz43_518869	U11683	Trypanoplasma borreli Tt-JH mitochondrion cytochrome c oxidase subunit I (cox1) gene, complete cds	8.10E-07
958	3646.E20.gz43_519157	AB006216	Pasteurella multocida PM70 section 183 of 204 of the complete genome	2.30E-05
959	3646.H04.gz43_518904	AF043740	Branchiostoma floridae amphioxus Otx transcription factor (Otx) mRNA, complete cds	2.00E-06
960	3646.H09.gz43_518984	AP000145	Homo sapiens genomic DNA, chromosome 21q21.2, LL56-APP region, clone B2291C14-R44F3, segment 10/10, complete sequence	2.90E-40
961	3646.H16.gz43_519096	U22342	Bacteriophage T270 integrase (int) gene, complete cds	1.00E-07
962	3646.I01.gz43_518857	X54486	Human gene for C1-inhibitor	6.80E-05
963	3646.J03.gz43_518890	AB055372	Macaca fascicularis brain cDNA, clone: Qf1A-12842	5.40E-190
964	3646.J22.gz43_519194	AL133032	Homo sapiens mRNA; cDNA DKFZp586B03.17 (from clone DKFZp586B03.17)	2.00E-06
965	3646.K14.gz43_519067	AF239178	Paracoccidioides brasiliensis ion proteinase gene, complete cds; nuclear gene for mitochondrial product	4.00E-06
966	3646.L17.gz43_519116	Z58907	H. sapiens CpG island DNA genomic MseI fragment, clone 116a6, forward read cpg116a6.f11a	2.50E-70
967	3646.O13.gz43_519055	AL050391	Homo sapiens mRNA; cDNA DKFZp586A181 (from clone DKFZp586A181); partial cds	5.20E-08
968	3646.O16.gz43_519103	X00331	Drosophila virilis simple DNA sequence (pDV-161)	5.20E-08
969	3646.P09.gz43_518992	U04527	Borrelia burgdorferi 212 DNA gyrase b subunit (gyrB) and ribonuclease P protein component (rnpA) genes, partial cds, DnaA protein (dnaA), DNA polymerase III beta subunit (dnaN), and ribosomal protein L34 (rpmH) genes, complete cds	5.00E-06
970	3646.P14.gz43_519072	AY032863	Mus musculus chloride-formate exchanger mRNA, complete cds	8.00E-06
971	3646.P17.gz43_519120	U19569	Human squamous cell carcinoma antigen (SCCA2) gene, exon 1	1.20E-07

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
972	3661.A08.gz43_519483	AB017511	Hydra magnipapillata mRNA for PLC-betaH1, complete cds	1.20E-05
973	3661.D17.gz43_519630	J03488	Reovirus type 3 L2 gene encoding guanylyltransferase, complete cds	3.00E-06
974	3661.D18.gz43_519646	AB033024	Homo sapiens mRNA for KIAA1198 protein, partial cds	1.90E-11
975	3661.E19.gz43_519663	AB014084	Homo sapiens genomic DNA, chromosome 6p21.3, HLA class I region, Cosmid clone:TY7A5, complete sequence	6.00E-05
976	3661.E23.gz43_519727	AE001032	Archaeoglobus fulgidus section 75 of 172 of the complete genome	5.30E-05
977	3661.F14.gz43_519584	X15063	Plasmodium falciparum mRNA for major merozoite surface antigen gp195	6.80E-05
978	3661.G16.gz43_519617	AF255609	Homo sapiens high mobility group protein HMG1 gene, exons 1 and 2, partial cds	2.70E-07
979	3661.G20.gz43_519681	AK021558	Homo sapiens cDNA FLJ11496 fis, clone HEMBA1001964	6.40E-09
980	3661.H11.gz43_519538	Z30705	Puumala virus (Evo/15Cg/93) gene for N protein	3.90E-07
981	3661.H24.gz43_519746	X66979	X.laevus mRNA XLFLI	1.60E-05
982	3661.I22.gz43_519715	AF029887	Caenorhabditis elegans UNC-129 (unc-129) mRNA, complete cds	5.00E-06
983	3661.J15.gz43_519604	AJ297538	Homo sapiens partial RARA gene, intron 2	4.00E-06
984	3661.K22.gz43_519717	AK002100	Homo sapiens cDNA FLJ11238 fis, clone PLACE1008532	1.30E-13
985	3661.L19.gz43_519670	AL589643	Human DNA sequence from clone RP11-344C1 on chromosome 6, complete sequence [Homo sapiens]	2.20E-05
986	3661.M03.gz43_519415	Z57613	H.sapiens CpG island DNA genomic MseI fragment, clone 187a12, forward read cpg187a12.fl1a	1.20E-08
987	3661.M23.gz43_519735	X79547	Equus caballus mitochondrial DNA complete sequence	5.80E-05
988	3661.P22.gz43_519722	AF055668	Mus musculus apoptosis-linked gene 4, deltaC form (Alg-4) mRNA, partial cds	8.00E-06
989	3662.A13.gz43_519947	Z49438	S.cerevisiae chromosome X reading frame ORF YJL163c	3.00E-06
990	3662.B13.gz43_519948	AB045237	Xenopus laevis XRPTPb mRNA for receptor-type protein tyrosine phosphatase beta.11, complete cds	7.00E-06
991	3662.C10.gz43_519901	BC007905	Homo sapiens, Similar to retinal degeneration B beta, clone MGC:14375 IMAGE:4299595, mRNA, complete cds	1.20E-09
992	3662.C15.gz43_519981	M33864	Human (clone HGL-3) interstitial retinoid-binding protein 3 (RBP3) gene, exon 1	1.20E-05
993	3662.F13.gz43_519952	AB040935	Homo sapiens mRNA for KIAA1502 protein, partial cds	1.20E-61
994	3662.H14.gz43_519970	AB032757	Mus musculus gad65 gene for glutamate decarboxylase 65, partial cds	8.00E-07

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
995	3662.H23.gz43_520114	AK013013	Mus musculus 10, 11 days embryo cDNA, RIKEN full-length enriched library, clone:2810406L04, full insert sequence	2.00E-06
996	3662.H24.gz43_520130	D45371	Human apM1 mRNA for GS3109 (novel adipose specific collagen-like factor), complete cds	9.60E-10
997	3662.J05.gz43_519828	M83554	H.sapiens lymphocyte activation antigen CD30 mRNA, complete cds	1.40E-05
998	3662.J08.gz43_519876	Z11876	B.hernslii vmp7 gene encoding Vmp7 outer membrane lipoprotein	1.11E-04
999	3662.J09.gz43_519892	AB011101	Homo sapiens mRNA for KIAA0529 protein, partial cds	6.30E-05
1000	3662.J16.gz43_520004	U00484	Anabaena PCC7120 protein kinase PknA (pknA) gene, complete cds	2.00E-06
1001	3662.K03.gz43_519797	AL390145	Homo sapiens mRNA; cDNA DKFZp762C115 (from clone DKFZp762C115)	1.40E-05
1002	3662.L05.gz43_519830	U63635	Schizosaccharomyces pombe RNA lariat debranching enzyme (Sp-dbr1) gene, complete cds	5.80E-10
1003	3662.N24.gz43_520136	Z37079	L.helveticus genes for prolinase and putative ABC transporter	3.70E-05
1004	3662.O02.gz43_519785	AF084460	Gallus gallus potassium channel Shaker alpha subunit variant cKv1.4(m) mRNA, complete cds	6.90E-05
1005	3662.P03.gz43_519802	AJ011456	Schizella tetragona matK gene (corresponding location in Tobacco: 963-1244)	7.20E-08
1006	3663.A09.gz43_520267	Z69608	A.rara SSU rRNA gene (partial)	3.30E-07
1007	3663.C08.gz43_520253	Z50756	Caenorhabditis elegans cosmid T08D10, complete sequence	7.60E-07
1008	3663.C19.gz43_520429	Z22672	H.sapiens cacn11a3 gene encoding skeletal muscle dhp-receptor alpha 1 subunit	2.80E-07
1009	3663.E04.gz43_520191	U89318	Homo sapiens nucleophosmin phosphoprotein (NPM) gene, intron 9, partial sequence	2.60E-07
1010	3663.F15.gz43_520368	U66073	Tritrichomonas foetus putative superoxide dismutase 1 (SOD1) gene, complete cds	9.20E-07
1011	3663.F22.gz43_520480	U36786	Rattus norvegicus putative pheromone receptor VN7 mRNA, complete cds	7.10E-07
1012	3663.G01.gz43_520145	AK024359	Homo sapiens cDNA FLJ14297 fls, clone PLACE1008941	9.50E-36
1013	3663.G08.gz43_520257	L19339	Molgula oculata zinc finger protein (manx) mRNA, complete cds	5.20E-07
1014	3663.H20.gz43_520450	X61307	Staphylococcus aureus spa gene for protein A	5.00E-06

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
1015	3663.J06.gz43 520228	AE007916	<i>Agrobacterium tumefaciens</i> strain C58 plasmid AT, section 44 of 50 of the complete sequence	2.02E-04
1016	3663.J16.gz43 520388	U38181	<i>Leuconostoc mesenteroides</i> dextranucrase gene, complete cds	3.90E-07
1017	3663.K02.gz43 520165	X68339	<i>Mycoplasma</i> -like organism (substrain ASHY) DNA for 16S rRNA	5.00E-06
1018	3663.K13.gz43 520341	AF155221	<i>Mus musculus</i> matrix metalloproteinase 19 (Mmp19) mRNA, complete cds	2.00E-06
1019	3663.L18.gz43 520422	AB031056	<i>Solobacterium moorei</i> gene for 16S rRNA, isolate:RCA59-74	1.00E-06
1020	3663.L24.gz43 520518	D10445	Mouse mRNA for protein C, complete cds	6.00E-06
1021	3663.M24.gz43 520519	AB001196	<i>Treponema pallidum</i> section 12 of 87 of the complete genome	5.20E-05
1022	3663.N09.gz43 520280	AF081788	<i>Homo sapiens</i> putative spliceosome associated protein mRNA, complete cds	4.00E-20
1023	3663.N10.gz43 520296	X62423	<i>P. falciparum</i> pol delta gene for DNA polymerase delta	4.00E-06
1024	3663.N12.gz43 520328	AF178079	<i>Zygosaccharomyces rouxii</i> ketoreductase (krd) mRNA, complete cds	5.00E-06
1025	3663.N16.gz43 520392	U41060	<i>Homo sapiens</i> estrogen regulated LIV-1 protein (LIV-1) mRNA, complete cds	2.00E-06
1026	3663.O07.gz43 520249	D00442	Grapevine fanleaf virus satellite RNA (RNA3), complete cds	1.50E-08
1027	3663.O09.gz43 520281	AK002141	<i>Homo sapiens</i> cDNA FLJ11279 fis, clone PLACE1009444, highly similar to PHOSPHATIDYLINOSITOL 4-KINASE ALPHA (EC 2.7.1.67)	5.30E-10
1028	3664.A11.gz43 520683	U67525	<i>Methanococcus jannaschii</i> section 67 of 150 of the complete genome	4.00E-06
1029	3664.C21.gz43 520845	AF064773	<i>Staphylococcus aureus</i> extracellular enterotoxin type G precursor (SEG) gene, complete cds	1.30E-07
1030	3664.D06.gz43 520606	AF178079	<i>Zygosaccharomyces rouxii</i> ketoreductase (krd) mRNA, complete cds	5.00E-06
1031	3664.D12.gz43 520702	U10519	Human DNA polymerase beta gene, exon 5	2.00E-07
1032	3664.D17.gz43 520782	AK027226	<i>Homo sapiens</i> cDNA: FLJ23573 fis, clone LNG12520	4.90E-07
1033	3664.E18.gz43 520799	AF317204	<i>Mus musculus</i> C-type lectin superfamily 1 gene, complete cds	3.20E-05
1034	3664.E23.gz43 520879	AB050903	<i>Mus musculus</i> mRNA for a4 subunit isoform, complete cds	3.00E-06
1035	3664.E24.gz43 520895	Z92793	<i>Caenorhabditis elegans</i> cosmid H15M21, complete sequence	1.20E-05
1036	3664.G12.gz43 520705	AF211482	<i>Dictyostelium discoideum</i> SdhA (sdhA) gene, complete cds	2.30E-09
1037	3664.G20.gz43 520833	- M14450	Rat thyrotropin (TSH) beta-subunit gene, exons 2 and 3	4.00E-06
1038	3664.H15.gz43 520754	Y11270	<i>E.histolytica</i> INO1 gene	2.00E-06

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
1039	3664.H22.gz43_520866	X97773	B.taurus mRNA for mitochondrial tricarboxylate carrier protein	1.20E-05
1040	3664.J12.gz43_520708	M58150	Bovine lactoperoxidase (LPO) mRNA, complete cds	3.20E-05
1041	3664.J23.gz43_520884	U67463	Methanococcus jannaschii section 5 of 150 of the complete genome	3.00E-06
1042	3664.K16.gz43_520773	Z83118	Caenorhabditis elegans cosmid M04D5, complete sequence	2.70E-07
1043	3664.K19.gz43_520821	U36927	Plasmodium yoelii thoptry protein gene, complete cds	3.00E-05
1044	3664.L21.gz43_520854	AF057695	Haemophilus ducreyi strain 35000 putative phosphomannomutase (pnm) gene, partial cds; large supernatant protein 1 (lspA1) gene, complete cds; and putative GMP synthase (gnaA) gene, partial cds	2.15E-04
1045	3664.O22.gz43_520873	U43574	Hydra vulgaris nucleoporin p62 gene, complete cds	7.00E-06
1046	3664.P12.gz43_520714	AF030883	Mus musculus tRNA-His gene, complete sequence; platelet-activating factor acetylhydrolase Ib alpha subunit (Pafahap1) pseudogene, complete sequence; and tRNA-Glu gene, complete sequence	9.00E-06
1047	3664.P18.gz43_520810	Z47735	H.sapiens NFkB1 gene, exons 11 & 12	1.32E-04
1048	3665.A23.gz43_521259	X66979	X.laevis mRNA XLFLI	1.60E-05
1049	3665.B01.gz43_520908	M90058	Human serglycin gene, exons 1, 2, and 3	4.00E-06
1050	3665.B12.gz43_521084	AK020877	Mus musculus adult retina cDNA, RIKEN full-length enriched library, clone:A930019H03, full insert sequence	7.10E-07
1051	3665.E11.gz43_521071	AB024030	Arabidopsis thaliana genomic DNA, chromosome 5, TAC clone:K5A21	9.00E-06
1052	3665.E20.gz43_521215	X76584	H.sapiens simple DNA sequence region clone wgl1h1	6.80E-08
1053	3665.H20.gz43_521218	X95301	D.rerio mRNA for HER-5 protein	9.50E-07
1054	3665.K01.gz43_520917	X04653	Mouse mRNA for Ly-6 alloantigen (Ly-6E.1)	1.30E-05
1055	3665.M01.gz43_520919	AF098352	Wiseana copularis haplotype southern cytochrome oxidase subunit I and cytochrome oxidase subunit II genes, partial cds; mitochondrial genes for mitochondrial products	5.80E-07
1056	3665.M21.gz43_521239	AF257480	Rana temporaria microsatellite SB80 sequence	3.30E-09
1057	3665.M23.gz43_521271	Y10623	C.pallidivittatus globin gene cluster E	1.10E-05
1058	3665.N24.gz43_521288	X95301	D.rerio mRNA for HER-5 protein	1.00E-06
1059	3665.O06.gz43_521001	AB007033	Mycobacterium tuberculosis CDC1551, section 119 of 280 of the complete genome	7.40E-05
1060	3665.O14.gz43_521129	AB033094	Homo sapiens mRNA for KIAA1268 protein, partial cds	2.10E-08

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
1061	3665.O15.gz43_521145	AK004557	Mus musculus adult male lung cDNA, RIKEN full-length enriched library, clone:1200003C23, full insert sequence	1.20E-05
1062	3665.O19.gz43_521209	AY036905	Trichoderma atroviride protein GTPase Tga1 (tga1) gene, complete cds	2.10E-08
1063	3665.O21.gz43_521241	U89293	Homo sapiens MSH4 (HMSH4) mRNA, complete cds	1.20E-39
1064	3665.O23.gz43_521273	X00048	Herpes simplex virus (HSV) type 2 transforming region mtr-2 (map coordinates 0.580 - 0.625)	6.00E-06
1065	3665.P13.gz43_521114	Z48796	H.sapiens Ski-W mRNA for helicase	1.70E-05
1066	3666.A07.gz43_521387	AK005546	Mus musculus adult female placenta cDNA, RIKEN full-length enriched library, clone:1600027G01, full insert sequence	1.20E-07
1067	3666.A19.gz43_521579	AB011101	Homo sapiens mRNA for KIAA0529 protein, partial cds	5.80E-05
1068	3666.A24.gz43_521659	AL050208	Homo sapiens mRNA; cDNA DKFZp586F2323 (from clone DKFZp586F2323)	2.90E-07
1069	3666.B11.gz43_521452	X06932	Petunia hsp70 gene	3.00E-06
1070	3666.C18.gz43_521565	Z22672	H.sapiens cacn1a3 gene encoding skeletal muscle dhp-receptor alpha 1 subunit	2.80E-07
1071	3666.D02.gz43_521310	AJ297538	Homo sapiens partial RARA gene, intron 2	4.00E-06
1072	3666.D11.gz43_521454	AF057695	Haemophilus ducreyi strain 35000 putative phosphomannomutase (pmm) gene, partial cds; large supernatant protein 1 (lspA1) gene, complete cds; and putative GMP synthase (guaA) gene, partial cds	2.43E-04
1073	3666.D15.gz43_521518	Z66194	H.sapiens CpG island DNA genomic MseI fragment, clone 80b12, forward read cpg80b12.fl1b	1.70E-66
1074	3666.D16.gz43_521534	Z66194	H.sapiens CpG island DNA genomic MseI fragment, clone 80b12, forward read cpg80b12.fl1b	2.10E-37
1075	3666.F22.gz43_521632	U97062	Staphylococcus aureus NCTC 8325 SecA (secA) gene, complete cds	1.20E-08
1076	3666.G12.gz43_521473	J03901	Maize pyruvate,orthophosphate dikinase mRNA, complete cds	1.72E-04
1077	3666.I12.gz43_521475	AJ225102	Pinus lambertiana chloroplast DNA containing a SSR Black Hills (Oregon)	6.40E-10
1078	3666.L01.gz43_521302	M86227	Staphylococcus aureus DNA gyrase B subunit (gyrB) RecF homologue (recF) and DNA gyrase A subunit (gyrA) gene, complete cds	5.00E-06

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
1079	3666.L06.gz43_521382	AF224725	Trichosurus vulpecula retrovirus TVERV (type D) gag polyprotein (gag), protease (pro), and pol polyprotein (pol) genes, complete cds	3.30E-08
1080	3666.L11.gz43_521462	AF147081	Homo sapiens gamma-glutamyl hydrolase gene, exons 1 and 2	3.30E-05
1081	3666.L23.gz43_521654	AK020701	Mus musculus 6 days neonate skin cDNA, RIKEN full-length enriched library, clone:A030009B12, full insert sequence	2.20E-07
1082	3666.M16.gz43_521543	AF158179	Drosophila melanogaster strain Canton-S Chiffon-2 (chiffon) mRNA, alternative splice form 2, complete cds	4.40E-07
1083	3666.N06.gz43_521384	Z48796	H.sapiens Ski-W mRNA for helicase	1.70E-05
1084	3667.A15.gz43_524557	AF005903	Monodelphis domestica GTP-binding protein homolog mRNA, partial cds	7.80E-08
1085	3754.A08.gz43_532949	AF091502	Lactobacillus reuteri autoaggregation-mediating protein (aggH) gene, complete cds	1.00E-06
1086	3754.A13.gz43_533029	U02695	Protomelas similis clone PsiI 32 SATA satellite DNA sequence	7.60E-07
1087	3754.A16.gz43_533077	AE006577	Streptococcus pyogenes M1 GAS strain SF370, section 106 of 167 of the complete genome	9.00E-06
1088	3754.B04.gz43_532886	S83995	PstI fragment [Chlamydia pneumoniae, Genomic, 474 nt]	2.00E-06
1089	3754.B05.gz43_532902	AY008833	Staphylococcus aureus tcaR-tcaA-tcaB operon, complete sequences	5.00E-06
1090	3754.B07.gz43_532934	AF270216	Staphylococcus epidermidis strain SR1 clone step.1054h11 genomic sequence	9.50E-07
1091	3754.B08.gz43_532950	AK007308	Mus musculus adult male testis cDNA, RIKEN full-length enriched library, clone:1700128E15, full insert sequence	7.00E-06
1092	3754.B10.gz43_532982	AB002807	Drosophila melanogaster genomic scaffold 142000013385251, complete sequence	5.40E-05
1093	3754.C22.gz43_533175	D30612	Homo sapiens mRNA for repressor protein, partial cds	4.00E-06
1094	3754.D19.gz43_533128	L12043	Plasmodium falciparum unidentified mRNA sequence	3.00E-06
1095	3754.E12.gz43_533017	AB062933	Macaca fascicularis brain cDNA clone:QccE-22249, full insert sequence	3.60E-07
1096	3754.E20.gz43_533145	AL138746	Human DNA sequence from clone RP3-389B13 on chromosome Xq26.2-27.1, complete sequence [Homo sapiens]	8.30E-10
1097	3754.F01.gz43_532842	AF086820	Drosophila melanogaster paired-like homeodomain protein UNC-4 (unc-4) mRNA, complete cds	8.00E-06
1098	3754.F08.gz43_532954	S66402	vascular AT1a angiotensin receptor (exon 1, promoter) [rats, Sprague-Dawley, Genomic, 3477 nt]	3.10E-05

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
1099	3754.F11.gz43_533002	X57377	Mouse dilute myosin heavy chain gene for novel heavy chain with unique C-terminal region	2.10E-05
1100	3754.F15.gz43_533066	AJ245620	Homo sapiens CTL1 gene	2.50E-12
1101	3754.F20.gz43_533146	AE002426	Neisseria meningitidis serogroup B strain MC58 section 68 of 206 of the complete genome	3.70E-05
1102	3754.G03.gz43_532875	AF002166	Xenopus laevis Ig mu heavy chain switch region sequence	1.20E-07
1103	3754.G08.gz43_532955	X71020	N.tabacum Npg1 gene for polygalacturonase	6.80E-07
1104	3754.G18.gz43_533115	AF126531	Homo sapiens putative DNA-directed RNA polymerase III C11 subunit gene, complete cds	1.10E-13
1105	3754.H08.gz43_532956	L20127	Rochalimaea henselae antigen (htrA) gene, complete cds	4.60E-07
1106	3754.I01.gz43_532845	AK022138	Homo sapiens cDNA FLJ12076 fis, clone HEMBB1002442, weakly similar to LIN-10 PROTEIN	3.90E-14
1107	3754.I03.gz43_532877	AF016653	Caenorhabditis elegans cosmid C41D7, complete sequence	2.00E-06
1108	3754.J01.gz43_532846	U97408	Caenorhabditis elegans cosmid F48A9	4.00E-06
1109	3754.J05.gz43_532910	Z35484	Thermoanaerobacter sp. ATCC53627 cglA gene	4.00E-06
1110	3754.J10.gz43_532990	D17094	Human HepG2 partial cDNA, clone hmd5h04m5	5.10E-11
1111	3754.J12.gz43_533022	Z56695	H.sapiens CpG island DNA genomic MseI fragment, clone 136d4, reverse read cpg136d4.rt1a	1.00E-06
1112	3754.J24.gz43_533214	Y12855	Homo sapiens P2X7 gene, exon 12 and 13	2.50E-05
1113	3754.K14.gz43_533055	L79913	Xenopus laevis rds/peripherin (rds35) mRNA, complete cds	5.00E-06
1114	3754.K17.gz43_533103	AE006251	Lactococcus lactis subsp. lactis IL1403 section 13 of 218 of the complete genome	9.00E-06
1115	3754.K20.gz43_533151	AB047880	Macaca fascicularis brain cDNA, clone:OmpA-14303	1.00E-06
1116	3754.M08.gz43_532961	X58467	Human CYP2D7AP pseudogene for cytochrome P450 2D6	4.30E-11
1117	3754.N16.gz43_533090	U33116	Saccharomyces cerevisiae high copy DNA polymerase suppressor alpha mutation gene (PSP2), complete cds	1.80E-07
1118	3754.N19.gz43_533138	AK025312	Homo sapiens cDNA: FLJ21659 fis, clone COL08743	1.40E-07
1119	3754.N22.gz43_533186	AF081828	Ixodes hexagonus mitochondrial DNA, complete genome	4.00E-06
1120	3754.O18.gz43_533123	Z73229	S.cerevisiae chromosome XII reading frame ORF YLR057w	3.00E-06
1121	3754.O23.gz43_533203	AE006900	Sulfolobus solfataricus section 259 of 272 of the complete genome	1.10E-05
1122	3754.P13.gz43_533044	AF220217	Homo sapiens rsec15-like protein mRNA, partial cds	1.80E-10



Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
1123	3754.P17.gz43_533108	AJ250862	Bacillus sp. HIL-Y85/54728 mersacidin biosynthesis gene cluster (mrsK2, mrsR2, mrsI, mrsG, mrsE, mrsA, mrsR1, mrsD, mrsM and mrsT genes)	1.20E-05
1124	3756.A02.gz43_533237	AF285594	Homo sapiens testis protein TEX11 (TEX11) mRNA, complete cds	1.10E-05
1125	3756.A11.gz43_533381	U43148	Human patched homolog (PTC) mRNA, complete cds	4.00E-06
1126	3756.A13.gz43_533413	U56861	Nicotiana plumbaginifolia intergenic region between lhcb1*1 and lhcb1*2 genes	1.00E-06
1127	3756.B03.gz43_533254	AF101735	Pan troglodytes isolate PTOR3A5P olfactory receptor pseudogene, complete sequence	5.70E-08
1128	3756.B04.gz43_533270	Z82038	C.thermosaccharolyticum etfB, etfA, hbd, thlA and actA genes	1.00E-06
1129	3756.B15.gz43_533446	M96151	Mus musculus apolipoprotein B gene sequence	1.13E-04
1130	3756.B21.gz43_533542	Z92793	Caenorhabditis elegans cosmid H15M21, complete sequence	1.30E-05
1131	3756.B22.gz43_533558	U43542	Nicotiana tabacum diphenol oxidase mRNA, complete cds	2.00E-06
1132	3756.C06.gz43_533303	AB022085	Mus musculus Cctz-2 gene for chaperonin containing TCP-1 zeta-2 subunit, exon 5, 6, 7, 8, 9, 10	7.00E-05
1133	3756.C16.gz43_533463	AF143236	Homo sapiens apoptosis related protein APR2 mRNA, complete cds	5.00E-06
1134	3756.D08.gz43_533336	AB049544	Porcine enterovirus 10 gene for RNA-dependent RNA polymerase, partial cds	7.20E-07
1135	3756.D18.gz43_533496	X53658	E.coli DNA fragment	7.60E-08
1136	3756.D24.gz43_533592	X96861	H.virescens mRNA for pheromone binding protein	2.40E-07
1137	3756.E01.gz43_533225	AF202892	Mus musculus Kif21a (Kif21a) mRNA, complete cds	4.00E-06
1138	3756.E06.gz43_533305	AF139374	Homo sapiens DIR1 protein (DIR1) gene, complete cds	8.00E-06
1139	3756.E12.gz43_533401	AF238884	Botrytis virus F, complete genome	8.00E-06
1140	3756.E22.gz43_533561	U78866	Arabidopsis thaliana putative arginine-aspartate-rich RNA binding protein (gene1500), (gene1000), and (gene400) genes, complete cds	5.00E-06
1141	3756.F11.gz43_533386	D50091	Drosophila exoana G-3-P dehydrogenase (alphaGpdh) gene, exon 1-8, complete cds	2.00E-06
1142	3756.F16.gz43_533466	AJ233973	Gallus gallus microsatellite DNA GCT028 (CA) repeat	4.20E-07
1143	3756.G07.gz43_533323	AE000708	Aquifex aeolicus section 40 of 109 of the complete genome	6.00E-05
1144	3756.G12.gz43_533403	M84731	Pseudomonas sp. 5-substituted hydantoin racemase (hyuE) gene, complete cds	1.20E-05

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
1145	3756.G14.gz43_533435	AL116458	Botrytis cinerea strain T4 cDNA library under conditions of nitrogen deprivation	6.70E-07
1146	3756.I03.gz43_533261	U67550	Methanococcus jannaschii section 92 of 150 of the complete genome	2.30E-05
1147	3756.J05.gz43_533294	U11292	Human Ki nuclear autoantigen mRNA, complete cds	7.70E-07
1148	3756.K03.gz43_533263	AF073484	Homo sapiens MHC class I-related protein MR1 precursor (MR1) gene, signal peptide	8.00E-06
1149	3756.K07.gz43_533327	M37499	Human methylmalonyl CoA mutase (MUT) gene, exon 2	2.00E-06
1150	3756.K15.gz43_533455	AF248820	Maoricicada campbelli isolate TB-MC-016 tRNA-Asp gene, complete sequence; ATPase subunit 8 gene, complete cds; and ATPase subunit 6 gene, partial cds; mitochondrial genes for mitochondrial products	7.30E-07
1151	3756.K18.gz43_533503	M36300	S.cerevisiae glutamine amidotransferase (TRP3) gene, 3' end	2.30E-05
1152	3756.K20.gz43_533535	AY022480	Oryza sativa microsatellite MRG4805 containing (AGG)X8, genomic sequence	2.00E-10
1153	3756.L02.gz43_533248	X03833	Human gene for interleukin 1 alpha (IL-1 alpha)	2.80E-12
1154	3756.L03.gz43_533264	AF244246	Dysdercus sp. MC cytochrome c oxidase I (COI) gene, partial cds; mitochondrial gene for mitochondrial product	2.70E-07
1155	3756.L19.gz43_533520	AJ002732	Schizosaccharomyces pombe mRNA for ribosomal protein l14	2.00E-06
1156	3756.M06.gz43_533313	AK002951	Mus musculus adult male brain cDNA, RIKEN full-length enriched library, clone:0710001E20, full insert sequence	3.60E-07
1157	3756.M07.gz43_533329	AF055708	Populus balsamifera subsp. trichocarpa PTD protein (PTD) gene, complete cds	2.60E-07
1158	3756.M20.gz43_533537	Z35821	S.cerevisiae chromosome II reading frame ORF YBL060w	2.00E-06
1159	3756.N18.gz43_533506	AL591667	Human DNA sequence from clone RP11-389N9 on chromosome 6, complete sequence [Homo sapiens]	6.10E-05
1160	3756.N21.gz43_533554	AK026258	Homo sapiens cDNA: FLJ22605 fis, clone HSI04743	2.00E-06
1161	3756.O03.gz43_533267	U61347	Leiophyllum buxifolium ribosomal maturase (matK) gene, chloroplast gene encoding chloroplast protein, complete cds	4.20E-07
1162	3756.O07.gz43_533331	AF177871	Drosophila melanogaster small GTPase RHO1 (Rho1) gene, alternatively spliced products and complete cds	5.70E-07
1163	3756.O08.gz43_533347	M60705	Homo sapiens type I DNA topoisomerase gene, exons 19 and 20	6.00E-06
1164	3756.P08.gz43_533348	M60705	Homo sapiens type I DNA topoisomerase gene, exons 19 and 20	1.00E-05

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
1165	3759.C01.gz43_533607	X71874	H.sapiens genes for proteasome-like subunit (MECL-1), chymotrypsin-like protease (CTRL-1) and protein serine kinase (PSK-H1) last exon	4.00E-06
1166	3759.D15.gz43_533832	AL356790	Human DNA sequence from clone RPT11-238J15 on chromosome 20 Contains ESTs and GSSs. Contains part of the TOM gene for a putative mitochondrial outer membrane protein import receptor similar to yeast pre-mRNA splicing factors Prp1/Zcr1 and Prp6, complete>	1.10E-07
1167	3759.H08.gz43_533724	M31684	D.melanogaster cytoskeleton-like bicaudalD protein (BicD) mRNA, complete cds	2.00E-06
1168	3759.H15.gz43_533836	AB046001	Macaca fascicularis brain cDNA, clone:QccE-12738	2.60E-07
1169	3759.H17.gz43_533868	AE000706	Aquifex aeolicus section 38 of 109 of the complete genome	1.30E-05
1170	3759.H23.gz43_533964	AK027088	Homo sapiens cDNA: FLJ23435 fis, clone HRC12631	6.20E-34
1171	3759.I05.gz43_533677	AF056433	Homo sapiens clone FBD3 Cri-du-chat critical region mRNA	1.70E-07
1172	3759.I19.gz43_533901	Z69666	Human DNA sequence from cosmid 24F8 from a contig from the tip of the short arm of chromosome 16, spanning 2Mb of 16p13.3. Contains ESTs, repeat polymorphism and CpG island	2.06E-04
1173	3759.K05.gz43_533679	L01432	Soybean calmodulin (SCaM-3) mRNA, complete cds	4.10E-08
1174	3759.K17.gz43_533871	Z33340	M.capricorn DNA for CONTIG MC456	4.00E-06
1175	3759.L02.gz43_533632	U26736	Caenorhabditis elegans stomatin-like protein MEC-2 (mec-2) gene, complete cds	3.70E-05
1176	3759.L09.gz43_533744	M11180	Transposon Tn917 (complete), macrolide-lincosamide-streptogramin-B (MLS) resistance, complete cds	1.50E-07
1177	3759.L10.gz43_533760	AF117022	Solaria atropurpurea trnL gene, partial sequence; chloroplast gene for chloroplast product	4.40E-07
1178	3759.L15.gz43_533840	U22657	Mus musculus genomic locus related to cellular morphology	1.60E-05
1179	3759.L24.gz43_533984	AK022990	Homo sapiens cDNA FLJ12928 fis, clone NT2RP2004767	7.60E-10
1180	3759.M19.gz43_533905	M96324	Lycopersicon esculentum Ca2+-ATPase gene, complete cds	2.50E-05
1181	3759.N08.gz43_533730	AK005546	Mus musculus adult female placenta cDNA, RIKEN full-length enriched library, clone:1600027G01, full insert sequence	1.30E-07

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Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
1182	3759.N16.gz43 533858	AB014079	Homo sapiens genomic DNA, chromosome 6p21.3, HLA class I region, Cosmid clone:TY1E11, complete sequence	3.80E-12
1183	3759.N23.gz43 533970	AK018377	Mus musculus 16 days embryo lung cDNA, RIKEN full-length enriched library, clone:8430403M08, full insert sequence	5.70E-07
1184	3759.O16.gz43 533859	AE000918	Methanobacterium thermoautotrophicum from bases 1444576 to 1460617 (section 124 of 148) of the complete genome	1.40E-05
1185	3759.P03.gz43 533652	L06066	Saccharomyces cerevisiae PET117 polypeptide (PET117) gene, complete cds	5.90E-07
1186	3759.P13.gz43 533812	X89414	A.thaliana DNA for pyrroline-5-carboxylase synthetase gene	5.00E-06
1187	3759.P15.gz43 533844	X66979	X.laevis mRNA XLFLI	1.50E-05
1188	3759.P17.gz43 533876	AF039313	Moraxella catarrhalis strain LES-1 transferrin binding protein B (tbpB) gene, complete cds	2.00E-06
1189	3762.A09.gz43 534117	AE000496	Escherichia coli K12 MG1655 section 386 of 400 of the complete genome	1.63E-04
1190	3762.A16.gz43 534229	X98371	D.subobscura sex-lethal gene	7.00E-06
1191	3762.A19.gz43 534277	U95019	Human voltage-dependent calcium channel beta-2c subunit mRNA, complete cds	6.10E-07
1192	3762.A20.gz43 534293	M10014	Homo sapiens map 4q28 fibrinogen (FGG) gene, alternative splice products, complete cds	8.00E-06
1193	3762.B05.gz43 534054	J05614	Human proliferating cell nuclear antigen (PCNA) gene, promoter region	1.40E-05
1194	3762.B15.gz43 534214	AJ297559	Homo sapiens partial PIK3CB gene for phosphatidylinositol 3-kinase catalytic subunit p110beta, exons 15-17	2.50E-05
1195	3762.C20.gz43 534295	M58580	Rabbit angiotensin-converting enzyme (ACE) gene, 5' end	3.10E-05
1196	3762.C23.gz43 534343	L27146	Human neurofibromatosis 2 (NF2) gene, exon 16	1.00E-06
1197	3762.D03.gz43 534024	U51305	Triticum aestivum alpha-gliadin storage protein pseudogene, complete cds	1.40E-05
1198	3762.D04.gz43 534040	AF263274	Chionodraco rastrospinosus isolate Cra7 alpha tubulin mRNA, complete cds	3.50E-07
1199	3762.D18.gz43 534264	M94764	Glycine max cv. Dare nodulin 26 gene fragment	2.50E-05
1200	3762.D19.gz43 534280	AE001446	Helicobacter pylori, strain J99 section 7 of 132 of the complete genome	3.30E-05
1201	3762.D22.gz43 534328	M73962	Bovine pregnancy-associated glycoprotein 1 mRNA, complete cds	4.00E-06
1202	3762.E01.gz43 533993	X63746	S.cerevisiae rbc34 and fun34 genes for DNA dependant RNA polymerase c (III)	4.00E-06
1203	3762.E10.gz43 534137	Z74847	S.cerevisiae chromosome XV reading frame ORF YOL105c	1.00E-05

Table 8

SEQ ID	SEQ NAME	ACCESSION	GENBANK DESCRIPTION	GENBANK SCORE
1204	3762.E15.gz43_534217	AF207841	Pyricularia grisea AVR-Pita (AVR-Pita) gene, complete cds	2.20E-09
1205	3762.E23.gz43_534345	M58600	Human heparin cofactor II (HCF2) gene, exons 1 through 5	3.60E-37
1206	3762.F08.gz43_534106	Z47066	Human cosmid Qc14G3 from Xq28 contains STSs	3.10E-09
1207	3762.F22.gz43_534330	AY034974	Arabidopsis thaliana unknown protein (F24J8.3) mRNA, complete cds	4.20E-07
1208	3762.G18.gz43_534267	Z28150	S.cerevisiae chromosome XI reading frame ORF YKL150w	2.00E-06
1209	3762.H12.gz43_534172	AF370230	Arabidopsis thaliana unknown protein (T21P5_16/AT3g03420) mRNA, complete cds	6.60E-08
1210	3762.I07.gz43_534093	U19569	Human squamous cell carcinoma antigen (SCCA2) gene, exon 1	4.60E-07
1211	3762.J03.gz43_534030	U22421	Mus musculus obesity protein (ob) gene, complete cds	5.30E-07
1212	3762.J18.gz43_534270	AB027966	Schizosaccharomyces pombe gene for Hypothetical protein, partial cds, clone:TB89	2.30E-08
1213	3762.K02.gz43_534015	AF273762	Homo sapiens 3-hydroxy-3-methylglutaryl-coenzyme reductase gene, exon 15	4.40E-14
1214	3762.K20.gz43_534303	K01464	Rat cardiac alpha-myosin heavy chain gene, 5' flank, 1st 3 exons	3.00E-06
1215	3762.L18.gz43_534272	Z49438	S.cerevisiae chromosome X reading frame ORF YJL163c	4.00E-06
1216	3762.L20.gz43_534304	XM_030040	Homo sapiens similar to KIAA0877 protein (H. sapiens) (LOC90219), mRNA	3.00E-06
1217	3762.M04.gz43_534049	AF002237	Anopheles gambiae clone 227 mRNA sequence	4.00E-06
1218	3762.M17.gz43_534257	M29688	S.cerevisiae PMS1 gene encoding DNA mismatch repair protein, complete cds	1.40E-08
1219	3762.M23.gz43_534353	M20006	Chicken tumor 10 c-myc DNA, exons 2 and 3	2.90E-09
1220	Clu1014734.con_1	AB027966	Schizosaccharomyces pombe gene for Hypothetical protein, partial cds, clone:TB89	3.00E-08
1221	Clu1036845.con_1	M34429	Human PVT-IGLC fusion protein mRNA, 5' end	1.37E-03

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Table 9

SEQ ID	SEQ NAME	PFAM NAME	PFAM DESCRIPTION	SCORE	START	END
89	3547.D19.GZ43_503986	DCI	DCI domain	30.64	411	493
137	3550.G02.GZ43_506101	rvt	Reverse transcriptase (RNA-dependent DNA polymerase)	47.32	321	611
321	3562.B22.GZ43_507952	7tm_1	7 transmembrane receptor (rhodopsin family)	37.16	154	479
321	3562.B22.GZ43_507952	Bowman-Birk leg	Bowman-Birk serine protease inhibitor family	45.92	292	450
321	3562.B22.GZ43_507952	Cation efflux	Cation efflux family	33.32	225	380
358	3565.E16.GZ43_508243	AP endonuclease1	AP endonuclease family 1	38.16	406	577
413	3571.A08.GZ43_508897	oxidored_ql	NADH:Ubiquinone/plastoquinone (complex I), various chains	30.04	297	393
417	3571.B13.GZ43_508978	EGF	EGF-like domain	38.88	243	355
418	3571.B22.GZ43_509122	EGF	EGF-like domain	38.88	243	355
431	3571.H10.GZ43_508936	WW	WW domain	54.92	487	576
591	3583.H13.GZ43_510520	Sre	C. elegans Ste G protein-coupled chemoreceptor	30.36	282	485
638	3590.I21.GZ43_512427	bZIP	bZIP transcription factor	33.68	166	308
645	3590.M03.GZ43_512142	protamine P1	Protamine P1	35.88	268	437
774	3608.L14.gz43_514237	Transposase_22	L1 transposable element	62.12	491	616
836	3617.P12.gz43_515561	AP endonuclease1	AP endonuclease family 1	39.84	63	254
905	3632.O06.gz43_517407	60s ribosomal	60s Acidic ribosomal protein	38.04	276	444
905	3632.O06.gz43_517407	60s ribosomal	60s Acidic ribosomal protein	36.44	13	98
995	3662.H23.gz43_520114	Glycoprotein_G	Pneumovirus attachment glycoprotein G	43.04	21	297
995	3662.H23.gz43_520114	Metallothio_5	Metallothionein family 5	47.88	231	345
995	3662.H23.gz43_520114	squash	Squash family serine protease inhibitor	34.6	222	301
995	3662.H23.gz43_520114	Syndecan	Syndecan domain	35.36	1	308
1012	3663.G01.gz43_520145	KRAB	KRAB box	95.08	424	484
1217	3762.M04.gz43_534049	protamine P1	Protamine P1	33.16	293	468

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Table 10

SEQ ID	SEQ NAME	PFAM NAME	PFAM DESCRIPTION	SCORE	START	END
1486	NTP_004511S13.3_4	Armadillo seg	Armadillo/beta-catenin-like repeat	1.8E-95	142	184
1486	NTP_004511S13.3_4	Armadillo seg	Armadillo/beta-catenin-like repeat	1.8E-95	186	226
1486	NTP_004511S13.3_4	Armadillo seg	Armadillo/beta-catenin-like repeat	1.8E-95	228	269
1486	NTP_004511S13.3_4	Armadillo seg	Armadillo/beta-catenin-like repeat	1.8E-95	271	311
1486	NTP_004511S13.3_4	Armadillo seg	Armadillo/beta-catenin-like repeat	1.8E-95	313	353
1486	NTP_004511S13.3_4	Armadillo seg	Armadillo/beta-catenin-like repeat	1.8E-95	355	395
1486	NTP_004511S13.3_4	Armadillo seg	Armadillo/beta-catenin-like repeat	1.8E-95	397	437
1486	NTP_004511S13.3_4	Armadillo seg	Armadillo/beta-catenin-like repeat	1.8E-95	440	480
1486	NTP_004511S13.3_4	Armadillo seg	Armadillo/beta-catenin-like repeat	1.8E-95	142	184
1486	NTP_004511S13.3_4	Armadillo seg	Armadillo/beta-catenin-like repeat	1.8E-95	186	226
1486	NTP_004511S13.3_4	Armadillo seg	Armadillo/beta-catenin-like repeat	1.8E-95	228	269
1486	NTP_004511S13.3_4	Armadillo seg	Armadillo/beta-catenin-like repeat	1.8E-95	271	311
1486	NTP_004511S13.3_4	Armadillo seg	Armadillo/beta-catenin-like repeat	1.8E-95	313	353
1486	NTP_004511S13.3_4	Armadillo seg	Armadillo/beta-catenin-like repeat	1.8E-95	355	395
1486	NTP_004511S13.3_4	Armadillo seg	Armadillo/beta-catenin-like repeat	1.8E-95	397	437
1486	NTP_004511S13.3_4	Armadillo seg	Armadillo/beta-catenin-like repeat	1.8E-95	440	480
1486	NTP_004511S13.3_4	IBB	Importin beta binding domain	5.8E-37	35	124
1486	NTP_004511S13.3_4	IBB	Importin beta binding domain	5.8E-37	35	124
1497	NTP_007592S2.3_10	histone	Core histone H2A/H2B/H3/H4	1.2E-10	2	97
1497	NTP_007592S2.3_10	histone	Core histone H2A/H2B/H3/H4	1.2E-10	2	97
1500	NTP_007867S7.3_3	GT121	GT121-like repeat	7.2E-76	106	171
1500	NTP_007867S7.3_3	GT121	GT121-like repeat	7.2E-76	295	370
1500	NTP_007867S7.3_3	GT121	GT121-like repeat	7.2E-76	106	171
1500	NTP_007867S7.3_3	GT121	GT121-like repeat	7.2E-76	295	370
1501	NTP_007867S8.3_1	GT121	GT121-like repeat	7.2E-76	122	187
1501	NTP_007867S8.3_1	GT121	GT121-like repeat	7.2E-76	311	386
1501	NTP_007867S8.3_1	GT121	GT121-like repeat	7.2E-76	122	187
1501	NTP_007867S8.3_1	GT121	GT121-like repeat	7.2E-76	311	386
1507	NTP_008558S2.3_2	GST_N	Glutathione S-transferase, N-terminal domain	4.6E-11	21	95
1507	NTP_008558S2.3_2	GST_N	Glutathione S-transferase, N-terminal domain	4.6E-11	21	95
1510	NTP_009526S2.3_3	CBS	CBS domain	4.8E-43	30	84

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Table 10

SEQ ID	SEQ NAME	PEAM NAME	PEAM DESCRIPTION	SCORE	START	END
1510	NTP_00952652.3_3	CBS	CBS domain	4.8E-43	111	165
1510	NTP_00952652.3_3	CBS	CBS domain	4.8E-43	186	239
1510	NTP_00952652.3_3	CBS	CBS domain	4.8E-43	258	311
1510	NTP_00952652.3_3	CBS	CBS domain	4.8E-43	30	84
1510	NTP_00952652.3_3	CBS	CBS domain	4.8E-43	111	165
1510	NTP_00952652.3_3	CBS	CBS domain	4.8E-43	186	239
1510	NTP_00952652.3_3	CBS	CBS domain	4.8E-43	258	311
1511	NTP_00952652.3_5	CBS	CBS domain	4.8E-43	30	84
1511	NTP_00952652.3_5	CBS	CBS domain	4.8E-43	111	165
1511	NTP_00952652.3_5	CBS	CBS domain	4.8E-43	186	239
1511	NTP_00952652.3_5	CBS	CBS domain	4.8E-43	258	311
1511	NTP_00952652.3_5	CBS	CBS domain	4.8E-43	30	84
1511	NTP_00952652.3_5	CBS	CBS domain	4.8E-43	111	165
1511	NTP_00952652.3_5	CBS	CBS domain	4.8E-43	186	239
1511	NTP_00952652.3_5	CBS	CBS domain	4.8E-43	258	311
1514	NTP_01001852.3_5	DAG PE-bind	Phorbol esters/diacylglycerol binding domain (C1 domain)	7.7E-23	154	203
1514	NTP_01001852.3_5	DAG PE-bind	Phorbol esters/diacylglycerol binding domain (C1 domain)	7.7E-23	387	426
1514	NTP_01001852.3_5	DAG PE-bind	Phorbol esters/diacylglycerol binding domain (C1 domain)	7.7E-23	154	203
1514	NTP_01001852.3_5	DAG PE-bind	Phorbol esters/diacylglycerol binding domain (C1 domain)	7.7E-23	387	426
1518	NTP_01075754.3_2	T-box	T-box	5.5E-14	935	1099
1518	NTP_01075754.3_2	T-box	T-box	5.5E-14	1142	1160
1518	NTP_01075754.3_2	T-box	T-box	5.5E-14	935	1099
1518	NTP_01075754.3_2	T-box	T-box	5.5E-14	1142	1160
1520	NTP_01113052.3_3	GATA	GATA zinc finger	1.5E-11	159	198
1520	NTP_01113052.3_3	GATA	GATA zinc finger	1.5E-11	159	198
1523	NTP_01143056.3_6	cadherin	Cadherin domain	7.4E-61	174	270
1523	NTP_01143056.3_6	cadherin	Cadherin domain	7.4E-61	284	390



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Table 10

SEQ ID	SEQ NAME	PEAM NAME	PEAM DESCRIPTION	SCORE	START	END
1523	NTP_011430S6.3_6	cadherin	Cadherin domain	7.4E-61	405	495
1523	NTP_011430S6.3_6	cadherin	Cadherin domain	7.4E-61	174	270
1523	NTP_011430S6.3_6	cadherin	Cadherin domain	7.4E-61	284	390
1523	NTP_011430S6.3_6	cadherin	Cadherin domain	7.4E-61	405	495
1525	NTP_017582S2.3_6	HMG box	HMG (high mobility group) box	6.8E-09	34	92
1525	NTP_017582S2.3_6	HMG box	HMG (high mobility group) box	6.8E-09	34	92
1542	NTP_026331S1.1_1	GTF21	GTF21-like repeat	7.2E-76	106	171
1542	NTP_026331S1.1_1	GTF21	GTF21-like repeat	7.2E-76	295	370
1542	NTP_026331S1.1_1	GTF21	GTF21-like repeat	7.2E-76	106	171
1542	NTP_026331S1.1_1	GTF21	GTF21-like repeat	7.2E-76	295	370

Table 11

Pr ID	Path ID	Grp	Anatom Loc	Size	Grade	Histo Grade	Local Invasion	Lymph Met	Lymph Met Incid	Reg Lymph Grade	Dist Met & Loc	Dist Met Grade	Comment
15	21	-III	Ascending colon	4.0	T3	G2	Extending into subserosal adipose tissue	Pos	3/8	N1	Neg	MX	invasive adenocarcinoma, moderately differentiated; focal perineural invasion is seen
52	71	II	Cecum	9.0	T3	G3	Invasion through muscularis propria, subserosal involvement; ileocecc. valve involvement	Neg	0/12	N0	Neg	M0	Hyperplastic polyp in appendix.
121	140	II	Sigmoid	6	T4	G2	Invasion of muscularis propria into serosa, involving submucosa of urinary bladder	Neg	0/34	N0	Neg	M0	Perineural invasion; donut anastomosis Neg. One tubulovillous and one tubular adenoma with no high grade dysplasia.
125	144	II	Cecum	6	T3	G2	Invasion through the muscularis propria into subserosal adipose tissue. Ileocecal junction.	Neg	0/19	N0	Neg	M0	patient history of metastatic melanoma
128	147	III	Transverse colon	5.0	T3	G2	Invasion of muscularis propria into percolonic fat	Pos	1/5	N1	Neg	M0	

Table 11

Pt ID	Path ID	Grp	Anatom Loc	Size	Grade	Histo Grade	Local Invasion	Lymph Met	Lymph Met Incld	Reg Lymph Grade	Dist Met & Loc	Dist Met Grade	Comment
130	149		Splenic flexure	5.5	T3		through wall and into surrounding adipose tissue	Pos	10/24	N2	Neg	M1	
133	152	II	Rectum	5.0	T3	G2	Invasion through muscularis propria into non-peritonealized pericolic tissue; gross configuration is annular.	Neg	0/9	N0	Neg	M0	Small separate tubular adenoma (0.4 cm)
141	160	IV	Cecum	5.5	T3	G2	Invasion of muscularis propria into pericolic adipose tissue, but not through serosa. Arising from tubular adenoma.	Pos	7/21	N2	Pos - Liver	M1	Perineural invasion identified adjacent to metastatic adenocarcinoma.
156	175	III	Hepatic flexure	3.8	T3	G2	Invasion through muscularis propria into subserosa/pericolic adipose, no serosal involvement. Gross configuration annular.	Pos	2/13	N1	Neg	M0	Separate tubulovillous and tubular adenomas

Table 11.

Pt ID	Path ID	Grp	Anatom Loc	Size	Grade	Histo Grade	Local Invasion	Lymph Met	Lymph Met Incid	Reg Lymph Grade	Dist Met & Loc	Dist Met Grade	Comment
228	247	III	Rectum	5.8	T3	G2 to G3	Invasion through muscularis propria to involve subserosal, perirectal adipose, and serosa	Pos	1/8	N1	Neg	MX	Hyperplastic polyps
264	283	II	Ascending colon	5.5	T3	G2	Invasion through muscularis propria into subserosal adipose tissue.	Neg	0/10	N0	Neg	M0	Tubulovillous adenoma with high grade dysplasia
266	285	III	Transverse colon	9	T3	G2	Invades through muscularis propria to involve pericolic pericolic adipose, extends to serosa.	Neg	0/15	N1	Pos - Mesenteric deposit	MX	
268	287	I	Cecum	6.5	T2	G2	Invades full thickness of muscularis propria, but mesenteric adipose free of malignancy.	Neg	0/12	N0	Neg	M0	
278	297	III	Rectum	4	T3	G2	Invasion into perirectal adipose tissue.	Pos	7/10	N2	Neg	M0	Descending colon polyps, no HGD or carcinoma identified..

Table 11

Pt ID	Path ID	Grp	Anatom Loc	Size	Histo Grade	Local Invasion	Lymph Met	Lymph Met Incid	Reg. Lymph Grade	Dist Met & Loc	Dist Met Grade	Comment
296	315	III	Cecum	5.5	T3	G2	Pos	2/12	N1	Neg	M0	Tubulovillous adenoma (2.0 cm) with no high grade dysplasia. Neg. liver biopsy.
339	358	II	Rectosigmoid	6	T3	G2	Neg	0/6	N0	Neg	M0	1 hyperplastic polyp identified
341	360	II	Ascending colon	2 cm invasive	T3	G2	Neg	0/4	N0	Neg	MX	
356	375	II	Sigmoid	6.5	T3	G2	Neg	0/4	N0	Neg	M0	
360	412	III	Ascending colon	4.3	T3	G2	Pos	1/5	N1	Neg	M0	Two mucosal polyps
392	444	IV	Ascending colon	2	T3	G2	Pos	1/6	N1	Pos - Liver	M1	Tumor arising at prior ileocolic surgical anastomosis.

Table 11

Pr ID	Path ID	Grp	Anatom Loc	Size	Grade	Histo Grade	Local Invasion	Lymph Met	Lymph Met Incid	Reg Lymph Grade	Dist Met & Loc	Dist Met Grade	Comment
393	445	II	Cecum	6.0	T3	G2	Cecum, invades through muscularis propria to involve subserosal adipose tissue but not serosa.	Neg	0/21	N0	Neg	M0	
413	465	IV	Cecum	4.8	T3	G2	Invasive through muscularis to involve periserosal fat; abutting ileocecal junction.	Neg	0/7	N0	Pos - Liver	M1	redagnosis of oophorectomy path to metastatic colon cancer.
505	383	IV		7.5	T3	G2	Invasion through muscularis propria involving peritoneal adipose, serosal surface uninvolved	Pos	2/17	N1	Pos - Liver	M1	Anatomical location of primary not noted in report. Evidence of chronic colitis.
517	395	IV	Sigmoid	3	T3	G2	penetrates muscularis propria, involves peritoneal fat.	Pos	6/6	N2	Neg	M0	No mention of distant met in report

Table 11

PTID	Path ID	Grp	Anatom Loc	Size	Grade	Histo Grade	Local Invasion	Lymph Met	Lymph Met Incld	Reg Lymph Grade	Dist Met & Loc	Dist Met Grade	Comment
534	553	II	Ascending colon	12	T3	G3	Invasion through the muscularis propria involving pericolic fat. Serosa free of tumor.	Neg	0/8	N0	Neg	M0	Omentum with fibrosis and fat necrosis. Small bowel with acute and chronic serositis, focal abscess and adhesions.
546	565	IV	Ascending colon	5.5	T3	G2	Invasion through muscularis propria extensively through submucosal and extending to serosa.	Pos	6/12	N2	Pos - Liver	M1	
577	596	II	Cecum	11.5	T3	G2	Invasion through the bowel wall, into subserosal adipose. Serosal surface free of tumor.	Neg	0/58	N0	Neg	M0	Appendix dilated and fibrotic, but not involved by tumor

Table 11

Pt ID	Path ID	Grp	Anatom Loc	Size	Grade	Histo Grade	Local Invasion	Lymph Met	Lymph Met Incid	Reg Lymph Grade	Dist Met & Loc	Dist Met Grade	Comment
695	714	II	Cecum	14.0	T3	G2	extending through bowel wall into serosal fat	Neg	0/22	N0	Neg	MX	moderately differentiated adenocarcinoma with mucinous differentiation (% not stated), tubular adenoma and hyperplastic polyps present,
784	803	IV	Ascending colon	3.5	T3	G3	through muscularis propria into pericolic soft tissues	Pos	5/17	N2	Pos - Liver	M1	invasive poorly differentiated adenocarcinoma
786	805	IV	Descending colon	9.5	T3	G2	through muscularis propria into pericolic fat, but not at serosal surface	Neg	0/12	N0	Pos - Liver	M1	moderately differentiated invasive adenocarcinoma <sup>a</sup>
787	806	II	Rectosigmoid	2.5	T3	G2-G3	Invasion of muscularis propria into soft tissue	Neg		N0	Neg	MX	Peritumoral lymphocytic response; 5 LN examined in pericolic fat, no metastases observed.
789	808	IV	Cecum	5.0	T3	G2-G3	Extending through muscularis propria into pericolic fat	Pos	5/10	N2	Pos - Liver	M1	Three lungating lesions examined.



Table 11

Pt ID	Path ID	Grp	Anatom Loc	Size	Grade	Histo Grade	Local Invasion	Lymph Met	Lymph Met Incid	Reg Lymph Grade	Dist Met & Loc	Dist Met Grade	Comment
790	809	IV	Rectum	6.8	T3	G1-G2	Invading through muscularis propria into perirectal fat	Pos	3/13	N1	Pos - Liver	M1	
791	810	IV	Ascending colon	5.8	T3	G3	Through the muscularis propria into pericolic fat	Pos	13/25	N2	Pos - Liver	M1	poorly differentiated invasive colonic adenocarcinoma
888	908	IV	Ascending colon	2.0	T2	G1	Into muscularis propria	Pos	3/21	N0	Pos - Liver	M1	well to moderately differentiated adenocarcinoma; this patient has tumors of the ascending colon and the sigmoid colon
889	909	IV	Cecum	4.8	T3	G2	Through muscularis propria int subserosal tissue	Pos	1/4	N1	Pos - Liver	M1	moderately differentiated adenocarcinoma
890	910	IV	Ascending colon		T3	G2	Through muscularis propria into subserosa.	Pos	11/15	N2	Pos - Liver	M1	
891	911	IV	Rectum	5.2	T3	G2	Invasion through muscularis propria into perirectal soft tissue	Pos	4/15	N2	Pos - Liver	M1	Perineural invasion present.

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Table 11

Pt ID	Path ID	Grp	Anatom Loc	Size	Grade	Histo Grade	Local Invasion	Lymph Met	Lymph Met Incid	Reg Lymph Grade	Dist Met & Loc	Dist Met Grade	Comment
892	912	IV	Sigmoid	5.0	T3	G2	Invasion into pericolic sort tissue. Tumor focally invading skeletal muscle attached to colon.	Pos	1/28	N1	Pos - Liver, left and right lobe, omentum	M1	Perineural invasion present, extensive. Patient with a history of colon cancer.
893	913	IV	Transverse colon	6.0	T3	G2-G3	Through muscularis propria into pericolic fat	Pos	14/17	N2	Pos - Liver	M1	Perineural invasion focally present. Omentum mass, but resection with no tumor identified.
989	1009	IV	Sigmoid	6.0	T3	G2	Invasion through colon wall and focally involving subserosal tissue.	Pos	1/7	N1	Pos - Liver	M1	Primary adenocarcinoma arising from tubulovillous adenoma.

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Table 13

SEQ ID	SEQ NAME	CLONE ID	BREAST PATIENTS >=2x	BREAST NUM RATIOS	COLON PATIENTS >=2x	COLON NUM RATIOS	COLON UM >=2x	COLON UM NUM RATIOS
56	3544.G06.GZ43_505397	M00084443A:E10			50	8		
287	3559.B18.GZ43_507504	M00084700A:C10			41,025641	39	33,33333	27
621	3590.D19.GZ43_512389	M00085031B:E03			37.5	8		
683	3596.F03.GZ43_512529	M00085171D:F05			70	40	60,71429	28
706	3599.K02.GZ43_512892	M00085222D:D07			70	40	60,71429	28
1059	3665.O06.gz43_521001	M00086277B:H06			57.5	40	50	12
1150	3756.K15.gz43_533455	M00085833B:E11			35,2941176	34	35,71429	28
1156	3756.M06.gz43_533313	M00085815C:E11		17				
1187	3759.P15.gz43_533844	M00085100B:C12			63,4146341	41		
1416	NT_00759282.3_10		50	10				
1427	NT_009296S1.3_1						42.9	28
1427	NT_009296S1.3_1				46.2	39	33.3	27
1427	NT_009296S1.3_1				48.7	39	44.4	27
1444	NT_01758252.3_6				61.5	39	55.6	27
1444	NT_01758252.3_6				61.5	39	55.6	27

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Table 15

Library ID	CMCC Number	CloneId	NRRL Number
ES219	5471	M00084879B:E01	B-30523
ES219	5471	M00083819B:E10	B-30523
ES219	5471	M00084942C:B10	B-30523
ES219	5471	M00084704C:B09	B-30523
ES219	5471	M00084887C:C07	B-30523
ES219	5471	M00084976B:A08	B-30523
ES219	5471	M00085011B:A01	B-30523
ES219	5471	M00084961A:C07	B-30523
ES219	5471	M00084960D:D02	B-30523
ES219	5471	M00084973A:B06	B-30523
ES219	5471	M00084928D:F06	B-30523
ES219	5471	M00084968C:D10	B-30523
ES219	5471	M00084973A:B06	B-30523
ES219	5471	M00084966A:A08	B-30523
ES219	5471	M00084919C:B04	B-30523
ES219	5471	M00085003C:D03	B-30523
ES219	5471	M00084968A:D01	B-30523
ES219	5471	M00084969D:C11	B-30523
ES219	5471	M00084899D:B01	B-30523
ES219	5471	M00084893C:A12	B-30523
ES219	5471	M00084890D:F09	B-30523
ES219	5471	M00084904A:D03	B-30523
ES219	5471	M00085029A:C02	B-30523
ES219	5471	M00084963D:D07	B-30523
ES219	5471	M00085147C:A04	B-30523
ES219	5471	M00085144B:C12	B-30523
ES219	5471	M00085124B:G05	B-30523
ES219	5471	M00085702B:G11	B-30523
ES219	5471	M00085203A:E06	B-30523
ES219	5471	M00085242A:C06	B-30523
ES219	5471	M00084980D:H08	B-30523
ES219	5471	M00085187B:C11	B-30523
ES219	5471	M00085021C:F06	B-30523
ES219	5471	M00085182B:E04	B-30523
ES219	5471	M00084930D:B08	B-30523
ES219	5471	M00084941B:E07	B-30523
ES219	5471	M00084424D:G07	B-30523
ES219	5471	M00084938B:F12	B-30523
ES219	5471	M00084853D:G03	B-30523
ES219	5471	M00084878B:B12	B-30523
ES219	5471	M00084889B:C02	B-30523
ES219	5471	M00084885D:A12	B-30523
ES219	5471	M00084845A:E02	B-30523
ES219	5471	M00084972B:H03	B-30523
ES219	5471	M00084908A:F03	B-30523
ES219	5471	M00084975A:G05	B-30523
ES219	5471	M00084941C:H04	B-30523
ES219	5471	M00084997D:H09	B-30523
ES219	5471	M00084491A:E08	B-30523
ES219	5471	M00083815C:H08	B-30523
ES219	5471	M00084501A:D06	B-30523
ES219	5471	M00084558D:G08	B-30523
ES219	5471	M00084510C:F02	B-30523

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Table 15

Library ID	CMCC Number	CloneId	NRRL Number
ES219	5471	M00084521C:H11	B-30523
ES219	5471	M00084446A:A05	B-30523
ES219	5471	M00084458A:G06	B-30523
ES219	5471	M00084377D:E08	B-30523
ES219	5471	M00084382A:D06	B-30523
ES219	5471	M00083816B:D08	B-30523
ES219	5471	M00084449B:C09	B-30523
ES219	5471	M00084431C:B02	B-30523
ES219	5471	M00084463A:B07	B-30523
ES219	5471	M00084487D:F04	B-30523
ES219	5471	M00083800C:E07	B-30523
ES219	5471	M00084468C:E07	B-30523
ES219	5471	M00084638A:E10	B-30523
ES219	5471	M00084439B:A08	B-30523
ES219	5471	M00084479D:E10	B-30523
ES219	5471	M00084455D:B03	B-30523
ES219	5471	M00084368D:C02	B-30523
ES219	5471	M00084642D:E08	B-30523
ES219	5471	M00084373A:F08	B-30523
ES219	5471	M00084364C:B06	B-30523
ES219	5471	M00084521B:E11	B-30523
ES219	5471	M00084385B:D03	B-30523
ES219	5471	M00084443C:H06	B-30523
ES219	5471	M00083803C:F03	B-30523
ES219	5471	M00084421C:B11	B-30523
ES219	5471	M00084434B:E06	B-30523
ES219	5471	M00083820B:C03	B-30523
ES219	5471	M00084246B:H03	B-30523
ES219	5471	M00084484C:B11	B-30523
ES219	5471	M00084410C:F10	B-30523
ES219	5471	M00083801B:H03	B-30523
ES219	5471	M00084980C:B07	B-30523
ES219	5471	M00084499C:C11	B-30523
ES219	5471	M00084526C:G09	B-30523
ES219	5471	M00084406C:A01	B-30523
ES219	5471	M00084380D:B07	B-30523
ES219	5471	M00084383B:A11	B-30523
ES219	5471	M00083834C:E02	B-30523
ES219	5471	M00083839A:H03	B-30523
ES219	5471	M00084505C:H08	B-30523
ES219	5471	M00084511D:A02	B-30523
ES219	5471	M00084494C:C01	B-30523
ES219	5471	M00084451D:F06	B-30523
ES219	5471	M00084604A:D02	B-30523
ES219	5471	M00084771D:G03	B-30523
ES219	5471	M00084817A:H11	B-30523
ES219	5471	M00084827D:D04	B-30523
ES219	5471	M00084843D:C06	B-30523
ES219	5471	M00084750C:B08	B-30523
ES219	5471	M00084757A:D01	B-30523
ES219	5471	M00084771D:A01	B-30523
ES219	5471	M00084730B:A09	B-30523
ES219	5471	M00084826B:E11	B-30523

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Table 15

Library ID	CMCC Number	CloneId	NRRL Number
ES219	5471	M00084595C:C07	B-30523
ES219	5471	M00084724A:C02	B-30523
ES219	5471	M00084833A:G07	B-30523
ES219	5471	M00084600D:B10	B-30523
ES219	5471	M00084634C:H02	B-30523
ES219	5471	M00084614D:A08	B-30523
ES219	5471	M00084620B:F05	B-30523
ES219	5471	M00084607A:B03	B-30523
ES219	5471	M00084633A:B12	B-30523
ES219	5471	M00084597A:F06	B-30523
ES219	5471	M00084575A:A11	B-30523
ES219	5471	M00084547B:B10	B-30523
ES219	5471	M00084525A:E08	B-30523
ES219	5471	M00084578B:E12	B-30523
ES219	5471	M00084669A:A05	B-30523
ES219	5471	M00084419C:A09	B-30523
ES219	5471	M00084769C:H03	B-30523
ES219	5471	M00085007A:B03	B-30523
ES219	5471	M00084865D:B04	B-30523
ES219	5471	M00084743D:G01	B-30523
ES219	5471	M00084770B:G12	B-30523
ES219	5471	M00084584B:A02	B-30523
ES219	5471	M00084647C:E12	B-30523
ES219	5471	M00084766D:F12	B-30523
ES219	5471	M00084648D:F05	B-30523
ES219	5471	M00084843A:D06	B-30523
ES219	5471	M00084709C:B02	B-30523
ES219	5471	M00084834B:G02	B-30523
ES219	5471	M00084718D:C04	B-30523
ES219	5471	M00084702B:C12	B-30523
ES219	5471	M00084645D:G02	B-30523
ES219	5471	M00084849B:F11	B-30523
ES219	5471	M00084859C:H05	B-30523
ES219	5471	M00084850D:H02	B-30523
ES219	5471	M00084857B:A09	B-30523
ES219	5471	M00084867A:C11	B-30523
ES219	5471	M00084823A:H01	B-30523
ES219	5471	M00084756B:H01	B-30523
ES219	5471	M00084700D:E09	B-30523
ES219	5471	M00085010C:H01	B-30523
ES219	5471	M00085060B:C05	B-30523
ES219	5471	M00085012C:A08	B-30523
ES219	5471	M00085047D:F03	B-30523
ES219	5471	M00085049B:E03	B-30523
ES219	5471	M00085051C:A01	B-30523
ES219	5471	M00085050A:E11	B-30523
ES219	5471	M00085676C:C04	B-30523
ES219	5471	M00085121A:D10	B-30523
ES219	5471	M00085166D:C10	B-30523
ES219	5471	M00084992D:B02	B-30523
ES219	5471	M00085148B:H01	B-30523
ES219	5471	M00085123B:C04	B-30523
ES219	5471	M00085173B:A08	B-30523

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Library ID	CMCC Number	Clonoid	NRRL Number
ES219	5471	M00085172C:F06	B-30523
ES219	5471	M00084937D:B04	B-30523
ES219	5471	M00085026D:A01	B-30523
ES219	5471	M00084994D:F11	B-30523
ES219	5471	M00085190C:D10	B-30523
ES219	5471	M00085194D:F04	B-30523
ES219	5471	M00085222D:D07	B-30523
ES219	5471	M00085223A:G01	B-30523
ES219	5471	M00084740C:B08	B-30523
ES219	5471	M00085056A:G12	B-30523
ES219	5471	M00084671A:C12	B-30523
ES219	5471	M00084571C:D05	B-30523
ES219	5471	M00084587B:H07	B-30523
ES219	5471	M00084582C:H03	B-30523
ES219	5471	M00084618C:A03	B-30523
ES219	5471	M00084687A:A03	B-30523
ES219	5471	M00085038A:C06	B-30523
ES219	5471	M00084722A:H12	B-30523
ES219	5471	M00084676B:E02	B-30523
ES219	5471	M00084615D:H12	B-30523
ES219	5471	M00084659C:G05	B-30523
ES219	5471	M00084536B:A03	B-30523
ES219	5471	M00084929C:B02	B-30523
ES219	5471	M00084652D:G11	B-30523
ES219	5471	M00084611B:A11	B-30523
ES219	5471	M00084530D:G07	B-30523
ES219	5471	M00084527C:H07	B-30523
ES219	5471	M00084545C:C05	B-30523
ES219	5471	M00084535D:C12	B-30523
ES219	5471	M00084684C:D02	B-30523
ES219	5471	M00084679D:G12	B-30523
ES219	5471	M00084734A:E04	B-30523
ES219	5471	M00084696D:H04	B-30523
ES220	5472	M00084724D:F04	B-30524
ES220	5472	M00084559B:F10	B-30524
ES220	5472	M00084707D:H03	B-30524
ES220	5472	M00084525D:H01	B-30524
ES220	5472	M00084578C:G09	B-30524
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ES220	5472	M00084596D:E10	B-30524
ES220	5472	M00084646A:D02	B-30524
ES220	5472	M00084572D:F07	B-30524
ES220	5472	M00084620A:E08	B-30524
ES220	5472	M00084553B:F04	B-30524
ES220	5472	M00084614D:B07	B-30524
ES220	5472	M00084604D:D08	B-30524

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Library ID	CMCC Number	CloneId	NRRL Number
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ES220	5472	M00084958C:B03	B-30524
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ES220	5472	M00085166C:A08	B-30524
ES220	5472	M00084467A:D06	B-30524
ES220	5472	M00084890C:A06	B-30524
ES220	5472	M00084609C:F10	B-30524
ES220	5472	M00084413C:A11	B-30524
ES220	5472	M00084834A:A03	B-30524
ES220	5472	M00085172A:G05	B-30524
ES220	5472	M00085146B:C01	B-30524
ES220	5472	M00085038A:B10	B-30524
ES220	5472	M00084246A:D03	B-30524
ES220	5472	M00084967B:D09	B-30524
ES220	5472	M00085035D:E04	B-30524
ES220	5472	M00084736B:H03	B-30524
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ES220	5472	M00084640D:A08	B-30524
ES220	5472	M00084443A:E10	B-30524
ES220	5472	M00084521A:E11	B-30524
ES220	5472	M00085019C:D05	B-30524
ES220	5472	M00084587C:A07	B-30524
ES220	5472	M00084616A:G03	B-30524
ES220	5472	M00084732B:A04	B-30524



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Library ID	CMCC Number	CloneId	NRRL Number
ES220	5472	M00084666A:C04	B-30524
ES220	5472	M00084633A:H05	B-30524
ES220	5472	M00084510C:F05	B-30524
ES220	5472	M00084648B:F06	B-30524
ES220	5472	M00084700D:H04	B-30524
ES220	5472	M00084506C:A05	B-30524
ES220	5472	M00084475C:G11	B-30524
ES220	5472	M00084673B:H11	B-30524
ES220	5472	M00084595D:D08	B-30524
ES220	5472	M00084636C:A06	B-30524
ES220	5472	M00084612C:B01	B-30524
ES220	5472	M00084644A:H05	B-30524
ES220	5472	M00084602D:B09	B-30524
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ES220	5472	M00084678C:C11	B-30524
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ES220	5472	M00084699A:G05	B-30524
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ES220	5472	M00084487D:F07	B-30524
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ES220	5472	M00084949B:B12	B-30524
ES220	5472	M00084746B:B04	B-30524
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ES220	5472	M00084742A:F07	B-30524
ES220	5472	M00083799D:F10	B-30524
ES220	5472	M00084760D:D09	B-30524
ES220	5472	M00084845C:H05	B-30524
ES220	5472	M00084927A:C01	B-30524
ES220	5472	M00084935B:E10	B-30524
ES220	5472	M00084974D:F11	B-30524
ES220	5472	M00084935C:E07	B-30524
ES220	5472	M00084503D:G10	B-30524
ES220	5472	M00084907C:C01	B-30524
ES220	5472	M00084893C:B01	B-30524
ES220	5472	M00083803B:F11	B-30524
ES220	5472	M00084945A:D10	B-30524
ES220	5472	M00084765B:A10	B-30524
ES220	5472	M00084455D:G03	B-30524

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Library ID	CMCC Number	CloneId	NRRL Number
ES220	5472	M00084874D:E03	B-30524
ES220	5472	M00084889C:A04	B-30524
ES220	5472	M00084846B:H07	B-30524
ES220	5472	M00084967B:B10	B-30524
ES220	5472	M00084838A:F12	B-30524
ES220	5472	M00084885A:C01	B-30524
ES220	5472	M00084823A:H06	B-30524
ES220	5472	M00084958B:E10	B-30524
ES220	5472	M00084399B:E05	B-30524
ES220	5472	M00084880B:D03	B-30524
ES220	5472	M00084877D:G07	B-30524
ES220	5472	M00084406B:C03	B-30524
ES220	5472	M00084856B:A12	B-30524
ES220	5472	M00084888D:A11	B-30524
ES220	5472	M00083831D:H11	B-30524
ES220	5472	M00084481D:C06	B-30524
ES220	5472	M00083834B:F09	B-30524
ES220	5472	M00084770D:B08	B-30524
ES220	5472	M00084976C:C12	B-30524
ES220	5472	M00085201C:C12	B-30524
ES220	5472	M00084379D:A05	B-30524
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ES220	5472	M00084988B:B08	B-30524
ES220	5472	M00084969C:H11	B-30524
ES220	5472	M00084988C:G03	B-30524
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ES220	5472	M00085243A:D07	B-30524
ES220	5472	M00085031B:E03	B-30524
ES220	5472	M00085164C:G05	B-30524
ES220	5472	M00085031C:D05	B-30524
ES220	5472	M00084251D:C05	B-30524
ES220	5472	M00085027A:C02	B-30524

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Library ID	CMCC Number	CloneId	NRRL Number
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ES220	5472	M00085209C:F11	B-30524
ES220	5472	M00084368D:D03	B-30524
ES220	5472	M00083818A:E09	B-30524
ES220	5472	M00084980C:E06	B-30524
ES220	5472	M00084248B:C06	B-30524
ES220	5472	M00085244C:D03	B-30524
ES220	5472	M00084987A:D09	B-30524
ES220	5472	M00084994A:H04	B-30524
ES220	5472	M00084970D:E08	B-30524
ES220	5472	M00085038D:D10	B-30524
ES220	5472	M00085035B:C12	B-30524
ES220	5472	M00085184D:B08	B-30524
ES221	5473	M00084666C:A06	B-30525
ES221	5473	M00084657C:E01	B-30525
ES221	5473	M00084540B:B08	B-30525
ES221	5473	M00084415C:C05	B-30525
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ES221	5473	M00084396B:B03	B-30525
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ES221	5473	M00084889D:G06	B-30525
ES221	5473	M00084959B:C07	B-30525
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ES221	5473	M00084489A:D12	B-30525
ES221	5473	M00084748A:D09	B-30525
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ES221	5473	M00084767B:D10	B-30525
ES221	5473	M00084711B:A05	B-30525
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ES221	5473	M00084466B:E01	B-30525
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ES221	5473	M00084492C:B05	B-30525
ES221	5473	M00084487B:A06	B-30525
ES221	5473	M00084480B:A05	B-30525
ES221	5473	M00084764D:G08	B-30525
ES221	5473	M00084743D:H04	B-30525
ES221	5473	M00084891D:A02	B-30525
ES221	5473	M00084822C:D06	B-30525
ES221	5473	M00084853D:A12	B-30525
ES221	5473	M00084822B:G11	B-30525
ES221	5473	M00084756D:C04	B-30525
ES221	5473	M00084839C:B09	B-30525

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Library ID	CMCC Number	CloneId	NRRL Number
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ES221	5473	M00084956C:G09	B-30525
ES221	5473	M00084908C:F07	B-30525
ES221	5473	M00084902B:A10	B-30525
ES221	5473	M00084833D:B04	B-30525
ES221	5473	M00085023D:E11	B-30525
ES221	5473	M00085151A:B04	B-30525
ES221	5473	M00085039D:F09	B-30525
ES221	5473	M00085169A:H12	B-30525
ES221	5473	M00085052B:E04	B-30525
ES221	5473	M00085171D:F05	B-30525
ES221	5473	M00085050A:B06	B-30525
ES221	5473	M00085155B:F10	B-30525
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ES221	5473	M00085145C:D02	B-30525
ES221	5473	M00084722D:G04	B-30525
ES221	5473	M00084721C:F09	B-30525
ES221	5473	M00084866B:A03	B-30525
ES221	5473	M00084727A:A02	B-30525
ES221	5473	M00084407A:H09	B-30525
ES221	5473	M00084855D:H05	B-30525
ES221	5473	M00084403D:D04	B-30525
ES221	5473	M00085144D:G03	B-30525

Table 15

Library ID	CMCC Number	CloneId	NRRL Number
ES221	5473	M00084880D:A10	B-30525
ES221	5473	M00084958C:B03	B-30525
ES221	5473	M00084888C:D12	B-30525
ES221	5473	M00084587C:G07	B-30525
ES221	5473	M00083844C:C04	B-30525
ES221	5473	M00084647D:C05	B-30525
ES221	5473	M00084528C:F06	B-30525
ES221	5473	M00084857D:A11	B-30525
ES221	5473	M00084385A:D02	B-30525
ES221	5473	M00084561C:D07	B-30525
ES221	5473	M00084994D:H04	B-30525
ES221	5473	M00084448B:D11	B-30525
ES221	5473	M00085006D:C10	B-30525
ES221	5473	M00084580B:B05	B-30525
ES221	5473	M00083814D:A10	B-30525
ES221	5473	M00084970C:G03	B-30525
ES221	5473	M00084372D:H11	B-30525
ES221	5473	M00084377B:E11	B-30525
ES221	5473	M00085230B:G08	B-30525
ES221	5473	M00084584B:F09	B-30525
ES221	5473	M00084584B:H12	B-30525
ES221	5473	M00085249C:C11	B-30525
ES221	5473	M00084441B:E05	B-30525
ES221	5473	M00083841A:G01	B-30525
ES221	5473	M00085006D:C04	B-30525
ES221	5473	M00084686B:B04	B-30525
ES221	5473	M00084998A:C12	B-30525
ES221	5473	M00085034B:E11	B-30525
ES221	5473	M00084683B:A01	B-30525
ES221	5473	M00084613A:A01	B-30525
ES221	5473	M00084633B:A06	B-30525
ES221	5473	M00085032C:F04	B-30525
ES221	5473	M00085022B:F05	B-30525
ES221	5473	M00084509A:E10	B-30525
ES221	5473	M00084400A:B09	B-30525
ES221	5473	M00084677C:F03	B-30525
ES221	5473	M00084427B:D01	B-30525
ES221	5473	M00083844B:C04	B-30525
ES221	5473	M00084598D:H05	B-30525
ES221	5473	M00084443B:C02	B-30525
ES221	5473	M00084514A:A03	B-30525
ES221	5473	M00084560C:G05	B-30525
ES221	5473	M00084504C:F05	B-30525
ES221	5473	M00084517C:D06	B-30525
ES221	5473	M00084420C:D03	B-30525
ES221	5473	M00084524D:D02	B-30525
ES221	5473	M00084499D:A10	B-30525
ES221	5473	M00085022B:B03	B-30525
ES221	5473	M00084958B:E10	B-30525
ES221	5473	M00084513C:C10	B-30525
ES221	5473	M00084595B:C08	B-30525
ES221	5473	M00083804A:I112	B-30525
ES221	5473	M00084859D:B03	B-30525

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Library ID	CMCC Number	CloneId	NRRL Number
ES221	5473	M00084641B:F08	B-30525
ES221	5473	M00084844A:E10	B-30525
ES221	5473	M00083838A:E05	B-30525
ES221	5473	M00083849C:F11	B-30525
ES221	5473	M00084461C:D06	B-30525
ES221	5473	M00084810D:B10	B-30525
ES221	5473	M00085047D:F08	B-30525
ES221	5473	M00084912D:G06	B-30525
ES221	5473	M00084645C:F07	B-30525
ES221	5473	M00084912D:A09	B-30525
ES221	5473	M00084862B:B01	B-30525
ES221	5473	M00084938C:G06	B-30525
ES221	5473	M00084534B:E12	B-30525
ES221	5473	M00084909C:G02	B-30525
ES221	5473	M00084973A:A01	B-30525
ES221	5473	M00084651B:G10	B-30525
ES221	5473	M00084925A:B08	B-30525
ES221	5473	M00084568D:A02	B-30525
ES221	5473	M00084456A:H04	B-30525
ES221	5473	M00084988C:B01	B-30525
ES221	5473	M00084842C:B07	B-30525
ES221	5473	M00084708A:A11	B-30525
ES221	5473	M00084602C:E04	B-30525
ES221	5473	M00084757B:F11	B-30525
ES221	5473	M00084483A:C06	B-30525
ES221	5473	M00084605B:H04	B-30525
ES221	5473	M00083812C:G02	B-30525
ES221	5473	M00084610D:H04	B-30525
ES221	5473	M00085056D:B12	B-30525
ES221	5473	M00085017C:A11	B-30525
ES221	5473	M00084573A:A10	B-30525
ES221	5473	M00084637B:E01	B-30525
ES221	5473	M00085056B:B06	B-30525
ES221	5473	M00084510C:H01	B-30525
ES221	5473	M00084577B:C08	B-30525
ES221	5473	M00084646B:B03	B-30525
ES221	5473	M00084844C:F04	B-30525
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ES221	5473	M00084930B:E12	B-30525
ES221	5473	M00084469B:F08	B-30525
ES221	5473	M00084569D:B04	B-30525
ES221	5473	M00084453D:B12	B-30525
ES221	5473	M00083844A:E12	B-30525
ES221	5473	M00085009D:A02	B-30525
ES221	5473	M00084619A:E04	B-30525
ES221	5473	M00085006C:C07	B-30525
ES222	5474	M00084459A:F10	B-30526
ES222	5474	M00084721B:C11	B-30526
ES222	5474	M00084454A:G08	B-30526
ES222	5474	M00084460D:B04	B-30526
ES222	5474	M00084723D:G09	B-30526
ES222	5474	M00084704A:C12	B-30526
ES222	5474	M00084487C:H06	B-30526

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Library ID	CMCC Number	CloneId	NRRL Number
ES222	5474	M00084867C:G02	B-30526
ES222	5474	M00084475B:D03	B-30526
ES222	5474	M00084490A:C12	B-30526
ES222	5474	M00084865D:G02	B-30526
ES222	5474	M00084876D:A06	B-30526
ES222	5474	M00084553D:G05	B-30526
ES222	5474	M00084558D:A04	B-30526
ES222	5474	M00084645B:A06	B-30526
ES222	5474	M00084747D:G02	B-30526
ES222	5474	M00084884D:D03	B-30526
ES222	5474	M00084700A:C10	B-30526
ES222	5474	M00084973A:C01	B-30526
ES222	5474	M00084493A:E03	B-30526
ES222	5474	M00084497B:C12	B-30526
ES222	5474	M00084500D:B11	B-30526
ES222	5474	M00084523C:C10	B-30526
ES222	5474	M00084526D:E09	B-30526
ES222	5474	M00084923D:B05	B-30526
ES222	5474	M00084962C:F10	B-30526
ES222	5474	M00084669C:A10	B-30526
ES222	5474	M00084444D:F09	B-30526
ES222	5474	M00084757B:F05	B-30526
ES222	5474	M00084922A:C08	B-30526
ES222	5474	M00084960D:D02	B-30526
ES222	5474	M00084837B:E06	B-30526
ES222	5474	M00084763D:A04	B-30526
ES222	5474	M00084651C:H01	B-30526
ES222	5474	M00084441D:E09	B-30526
ES222	5474	M00084509D:C02	B-30526
ES222	5474	M00084510D:D05	B-30526
ES222	5474	M00084657D:B10	B-30526
ES222	5474	M00084946D:H05	B-30526
ES222	5474	M00084506A:E08	B-30526
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ES222	5474	M00085247A:F05	B-30526
ES222	5474	M00085142D:F04	B-30526
ES222	5474	M00085151A:H09	B-30526
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ES222	5474	M00085141C:G06	B-30526
ES222	5474	M00085035D:D09	B-30526
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ES222	5474	M00084995B:B08	B-30526
ES222	5474	M00085008B:H11	B-30526
ES222	5474	M00085059B:H11	B-30526
ES222	5474	M00085125C:H06	B-30526
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ES222	5474	M00084418D:A04	B-30526
ES222	5474	M00084961C:A06	B-30526
ES222	5474	M00084766B:E03	B-30526
ES222	5474	M00084406A:B03	B-30526
ES222	5474	M00085686A:C05	B-30526
ES222	5474	M00085124A:G04	B-30526
ES222	5474	M00085059B:H07	B-30526

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Library ID	CMCC Number	CloneId	NRRL Number
ES222	5474	M00084646B:D07	B-30526
ES222	5474	M00084246B:D10	B-30526
ES222	5474	M00084974D:F11	B-30526
ES222	5474	M00084738B:A09	B-30526
ES222	5474	M00085015A:C09	B-30526
ES222	5474	M00083839B:G09	B-30526
ES222	5474	M00085012C:D06	B-30526
ES222	5474	M00085058A:H02	B-30526
ES222	5474	M00085009C:C01	B-30526
ES222	5474	M00083818C:A02	B-30526
ES222	5474	M00085007B:C07	B-30526
ES222	5474	M00085047A:H02	B-30526
ES222	5474	M00085245C:D07	B-30526
ES222	5474	M00085032D:C03	B-30526
ES222	5474	M00085039B:E09	B-30526
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ES222	5474	M00085018C:B09	B-30526
ES222	5474	M00084668D:D08	B-30526
ES222	5474	M00085175B:A03	B-30526
ES222	5474	M00084437C:G05	B-30526
ES222	5474	M00084967B:B10	B-30526
ES222	5474	M00084998B:A04	B-30526
ES222	5474	M00084716D:H03	B-30526
ES222	5474	M00084708B:A06	B-30526
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ES222	5474	M00085013A:E06	B-30526
ES222	5474	M00084374A:A10	B-30526
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ES222	5474	M00084515D:G03	B-30526
ES222	5474	M00084961D:H03	B-30526
ES222	5474	M00084908D:B11	B-30526
ES222	5474	M00084702A:B08	B-30526
ES222	5474	M00083838C:F07	B-30526
ES222	5474	M00084390B:H04	B-30526
ES222	5474	M00084734A:H01	B-30526
ES222	5474	M00083817B:A11	B-30526
ES222	5474	M00085176C:B11	B-30526
ES222	5474	M00084902C:F05	B-30526
ES222	5474	M00085677A:E02	B-30526
ES222	5474	M00084948D:B08	B-30526
ES222	5474	M00085190B:H04	B-30526
ES222	5474	M00084820D:A03	B-30526
ES222	5474	M00084479B:E04	B-30526
ES222	5474	M00084408D:E06	B-30526
ES222	5474	M00085009B:F10	B-30526
ES222	5474	M00085697A:F01	B-30526



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Library ID	CMCC Number	CloneId	NRRL Number
ES222	5474	M00084423D:B05	B-30526
ES222	5474	M00084973A:A01	B-30526
ES222	5474	M00083804B:C03	B-30526
ES222	5474	M00084841B:H09	B-30526
ES222	5474	M00084685D:B11	B-30526
ES222	5474	M00084599D:C02	B-30526
ES222	5474	M00084573D:G11	B-30526
ES222	5474	M00084603A:B07	B-30526
ES222	5474	M00084823D:E06	B-30526
ES222	5474	M00084565A:D10	B-30526
ES222	5474	M00084767B:F06	B-30526
ES222	5474	M00084963D:D07	B-30526
ES222	5474	M00084611A:A06	B-30526
ES222	5474	M00084829B:F06	B-30526
ES222	5474	M00084850C:A11	B-30526
ES222	5474	M00084540D:B12	B-30526
ES222	5474	M00084614C:G05	B-30526
ES222	5474	M00084826B:D12	B-30526
ES222	5474	M00084605D:G09	B-30526
ES222	5474	M00084923D:F11	B-30526
ES222	5474	M00084664D:E05	B-30526
ES222	5474	M00084533A:C04	B-30526
ES222	5474	M00084843D:F05	B-30526
ES222	5474	M00084894A:G09	B-30526
ES222	5474	M00084913B:F05	B-30526
ES222	5474	M00083817D:A08	B-30526
ES222	5474	M00084451D:A03	B-30526
ES222	5474	M00084675B:A04	B-30526
ES222	5474	M00084889A:B07	B-30526
ES222	5474	M00084879A:A04	B-30526
ES222	5474	M00084638D:A05	B-30526
ES222	5474	M00084468A:A09	B-30526
ES222	5474	M00084634A:D01	B-30526
ES222	5474	M00084577B:D04	B-30526
ES222	5474	M00084860B:A01	B-30526
ES222	5474	M00084567B:F03	B-30526
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ES222	5474	M00085226C:F08	B-30526
ES222	5474	M00084392C:D03	B-30526
ES222	5474	M00084389A:F12	B-30526
ES222	5474	M00084696C:A07	B-30526
ES222	5474	M00084397D:A09	B-30526
ES222	5474	M00085173A:B07	B-30526

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Library ID	CMCC Number	CloneId	NRRL Number
ES222	5474	M00084252B:H01	B-30526
ES222	5474	M00084970C:H09	B-30526
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ES222	5474	M00085245D:G07	B-30526
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ES222	5474	M00085220D:E06	B-30526
ES222	5474	M00084745A:H04	B-30526
ES222	5474	M00083809B:E08	B-30526
ES222	5474	M00084940B:F06	B-30526
ES222	5474	M00084533B:B10	B-30526
ES222	5474	M00084970D:B01	B-30526
ES222	5474	M00084583D:H12	B-30526
ES222	5474	M00084585D:H12	B-30526
ES222	5474	M00084581B:E06	B-30526
ES222	5474	M00084588B:D02	B-30526
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ES223	5475	M00085732A:B09	B-30527
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ES223	5475	M00085446A:D04	B-30527
ES223	5475	M00085697D:H11	B-30527
ES223	5475	M00086085A:H03	B-30527
ES223	5475	M00086057A:F07	B-30527
ES223	5475	M00086196A:F07	B-30527
ES223	5475	M00086279A:B07	B-30527

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Library ID	CMCC Number	CloneId	NRRL Number
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ES223	5475	M00086291D:B08	B-30527
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ES223	5475	M00086247A:G11	B-30527
ES223	5475	M00085373C:B06	B-30527
ES223	5475	M00085332D:B03	B-30527
ES223	5475	M00085632C:A06	B-30527
ES223	5475	M00085360B:G03	B-30527
ES223	5475	M00086191A:G09	B-30527
ES223	5475	M00085473C:B02	B-30527
ES223	5475	M00085600B:B03	B-30527
ES223	5475	M00083749D:B09	B-30527
ES223	5475	M00085301B:C10	B-30527
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ES223	5475	M00084772C:G12	B-30527
ES223	5475	M00086126C:D09	B-30527
ES223	5475	M00085808D:E01	B-30527
ES223	5475	M00085927A:F06	B-30527

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Table 15

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ES223	5475	M00085814C:C12	B-30527
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ES223	5475	M00086146D:A09	B-30527
ES223	5475	M00085076C:A07	B-30527
ES223	5475	M00085427C:A04	B-30527
ES223	5475	M00084774B:C10	B-30527
ES223	5475	M00085432A:H08	B-30527
ES223	5475	M00084796D:B01	B-30527
ES223	5475	M00086127C:C05	B-30527
ES223	5475	M00086160D:F08	B-30527
ES223	5475	M00084802A:H09	B-30527
ES223	5475	M00086081D:H11	B-30527
ES223	5475	M00086106D:H01	B-30527
ES223	5475	M00086159A:F05	B-30527
ES223	5475	M00084782D:H08	B-30527
ES223	5475	M00085956D:G04	B-30527
ES223	5475	M00085770C:A12	B-30527
ES223	5475	M00086008D:F08	B-30527
ES223	5475	M00086018A:A05	B-30527
ES223	5475	M00085761A:B03	B-30527
ES223	5475	M00085751A:A11	B-30527
ES223	5475	M00085956B:E08	B-30527
ES223	5475	M00085955C:C03	B-30527
ES223	5475	M00085904D:D02	B-30527
ES223	5475	M00085899C:G10	B-30527
ES223	5475	M00085927C:G10	B-30527
ES223	5475	M00085896D:A11	B-30527
ES223	5475	M00085892A:F04	B-30527
ES223	5475	M00085882B:F11	B-30527
ES223	5475	M00085419A:G09	B-30527
ES223	5475	M00085962B:A12	B-30527
ES223	5475	M00085811B:D12	B-30527
ES223	5475	M00085986B:H02	B-30527
ES223	5475	M00085922A:A08	B-30527
ES223	5475	M00085854C:F04	B-30527
ES223	5475	M00085835D:F06	B-30527
ES223	5475	M00086183A:H04	B-30527
ES223	5475	M00086193A:F04	B-30527
ES223	5475	M00086197B:A03	B-30527
ES223	5475	M00086203C:H04	B-30527
ES223	5475	M00085827D:D01	B-30527
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ES223	5475	M00085839B:B12	B-30527
ES223	5475	M00085750A:G03	B-30527
ES223	5475	M00086248A:H09	B-30527
ES223	5475	M00085964A:B11	B-30527
ES223	5475	M00086003D:G08	B-30527

Library ID	CMCC Number	CloneId	NRRL Number
ES223	5475	M00085819D:C02	B-30527
ES223	5475	M00085066B:D12	B-30527
ES223	5475	M00084775C:E05	B-30527
ES223	5475	M00085083A:E04	B-30527
ES223	5475	M00085090C:C09	B-30527
ES223	5475	M00085100A:H07	B-30527
ES223	5475	M00085101C:H03	B-30527
ES223	5475	M00085105D:H02	B-30527
ES223	5475	M00086323B:C04	B-30527
ES223	5475	M00084808D:A07	B-30527
ES223	5475	M00086003D:B10	B-30527
ES223	5475	M00084787B:D12	B-30527
ES223	5475	M00085068C:B03	B-30527
ES223	5475	M00086291A:E10	B-30527
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ES224	5476	M00084783C:A09	B-30528
ES224	5476	M00084782D:D10	B-30528
ES224	5476	M00085273A:A09	B-30528
ES224	5476	M00084794B:C01	B-30528
ES224	5476	M00084780D:D07	B-30528
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ES224	5476	M00085344D:F01	B-30528
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ES224	5476	M00085304B:D11	B-30528
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ES224	5476	M00085859B:A11	B-30528
ES224	5476	M00085860D:H02	B-30528
ES224	5476	M00085649B:A03	B-30528
ES224	5476	M00085815C:E11	B-30528

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Table 15

Library ID	CMCC Number	CloneId	NRRL Number
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ES224	5476	M00085449A:E02	B-30528
ES224	5476	M00085344A:G08	B-30528
ES224	5476	M00085520D:B11	B-30528
ES224	5476	M00086035A:C11	B-30528
ES224	5476	M00085955D:H10	B-30528
ES224	5476	M00086175C:D06	B-30528
ES224	5476	M00085510B:G12	B-30528
ES224	5476	M00085927C:G11	B-30528
ES224	5476	M00085919A:A05	B-30528
ES224	5476	M00085985C:D02	B-30528
ES224	5476	M00085934B:E12	B-30528
ES224	5476	M00085389C:H04	B-30528
ES224	5476	M00085406A:F03	B-30528
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ES224	5476	M00085830B:E09	B-30528
ES224	5476	M00085611C:D09	B-30528
ES224	5476	M00085810A:A10	B-30528
ES224	5476	M00085534D:H09	B-30528
ES224	5476	M00085390D:A03	B-30528
ES224	5476	M00085419C:H05	B-30528
ES224	5476	M00085441D:H10	B-30528
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ES224	5476	M00086000A:C05	B-30528
ES224	5476	M00085605A:D08	B-30528
ES224	5476	M00083706A:D02	B-30528
ES224	5476	M00086202C:A07	B-30528
ES224	5476	M00083691C:E12	B-30528

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Table 15

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ES224	5476	M00086031C:G11	B-30528
ES224	5476	M00085294D:G03	B-30528
ES224	5476	M00086114D:G11	B-30528
ES224	5476	M00085720D:A03	B-30528
ES224	5476	M00085262A:A02	B-30528
ES224	5476	M00085732A:G04	B-30528
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ES224	5476	M00083736A:D10	B-30528
ES224	5476	M00085301C:H11	B-30528
ES224	5476	M00085066D:A05	B-30528
ES224	5476	M00086294D:F08	B-30528
ES224	5476	M00084773C:D04	B-30528

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ES224	5476	M00086302A:E06	B-30528
ES224	5476	M00085084A:E12	B-30528
ES224	5476	M00085105C:D01	B-30528
ES224	5476	M00085297A:A11	B-30528
ES224	5476	M00085076C:H01	B-30528
ES224	5476	M00086286B:F08	B-30528
ES224	5476	M00085107D:H08	B-30528
ES224	5476	M00085092D:D09	B-30528
ES225	5477	M00086103D:I08	B-30529
ES225	5477	M00086272D:E04	B-30529
ES225	5477	M00085449B:G12	B-30529
ES225	5477	M00085832D:G02	B-30529
ES225	5477	M00085827C:F03	B-30529
ES225	5477	M00086328B:G12	B-30529
ES225	5477	M00085820D:D02	B-30529
ES225	5477	M00085522C:E05	B-30529
ES225	5477	M00086178A:D07	B-30529
ES225	5477	M00085651A:A01	B-30529
ES225	5477	M00085814B:G08	B-30529
ES225	5477	M00086149A:F06	B-30529
ES225	5477	M00085754C:A12	B-30529
ES225	5477	M00085383C:C05	B-30529
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ES225	5477	M00086081B:A06	B-30529
ES225	5477	M00086053B:F01	B-30529
ES225	5477	M00085259A:H05	B-30529
ES225	5477	M00083694C:F09	B-30529
ES225	5477	M00085643D:F06	B-30529
ES225	5477	M00086209B:H12	B-30529
ES225	5477	M00085860B:D10	B-30529
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ES225	5477	M00085626B:B11	B-30529
ES225	5477	M00086048D:H08	B-30529
ES225	5477	M00086010B:B05	B-30529
ES225	5477	M00083735C:H12	B-30529
ES225	5477	M00083722B:A07	B-30529
ES225	5477	M00085745C:C03	B-30529
ES225	5477	M00085809A:C05	B-30529



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Table 15

Library ID	CMCC Number	CloneId	NRRL Number
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ES225	5477	M00085764B:H12	B-30529
ES225	5477	M00086000C:B08	B-30529
ES225	5477	M00085817B:B08	B-30529
ES225	5477	M00085806B:A10	B-30529
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ES225	5477	M00085336C:G09	B-30529
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ES225	5477	M00086128D:H10	B-30529
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ES225	5477	M00085714C:G03	B-30529
ES225	5477	M00085741C:D06	B-30529
ES225	5477	M00085722A:A06	B-30529
ES225	5477	M00083704C:C04	B-30529
ES225	5477	M00085549D:G03	B-30529
ES225	5477	M00086143B:B08	B-30529
ES225	5477	M00085926C:C06	B-30529
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ES225	5477	M00085282C:F05	B-30529
ES225	5477	M00084784B:A02	B-30529
ES225	5477	M00085944A:F04	B-30529
ES225	5477	M00085804B:F09	B-30529
ES225	5477	M00085339C:C04	B-30529

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Library ID	CMCC Number	CloneId	NRRL Number
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ES225	5477	M00084779D:H10	B-30529
ES225	5477	M00085381C:A05	B-30529
ES225	5477	M00084783C:G08	B-30529
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ES225	5477	M00085302C:B06	B-30529
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ES225	5477	M00084791D:D01	B-30529
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ES225	5477	M00085893B:D08	B-30529
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Library ID	CMCC Number	CloneId	NRRL Number
ES226	5478	M00084956B:B05	B-30581
ES226	5478	M00084954D:D12	B-30581
ES226	5478	M00084948B:F04	B-30581
ES226	5478	M00084950D:F05	B-30581
ES226	5478	M00084950D:F01	B-30581
ES226	5478	M00084941D:C10	B-30581
ES226	5478	M00084950D:A06	B-30581
ES226	5478	M00084941D:H02	B-30581
ES226	5478	M00084954C:B12	B-30581
ES226	5478	M00084955A:E08	B-30581
ES226	5478	M00084954D:A05	B-30581
ES226	5478	M00084951A:D04	B-30581
ES226	5478	M00084954C:A03	B-30581

Table 16

SEQ ID	Seq Name	SpotID	Breast Cancer Tumor/Normal al >=2x Patients	Colon Cancer Tumor/Normal al >=2x Patients	Colon Cancer Tumor/Normal al >=2x Patients	Prostate Cancer Tumor/Normal al >=2x Patients	Colon Unmatched Med/Normal al >=2x Patients	Colon Matched Med/Normal al >=2x Patients	Colon Matched Med/Normal al >=2x Patients	Colon Unmatched Med/Normal al >=2x Patients
5	3541.A10.G243	506167	58048							
18	3538.K12.G243	506129	58775	23	26.08					
18	3544.A17.G243	506157	59100	23	26.08					
76	3544.A17.G243	506157	59100	76	44.74	31.96	63.64	33	62.78	36
176	3550.D18.G243	506822	42108	19	47.37	31.96	97	41.18	17	
181	3555.J14.G243	506860	60223	23	21.05	22.45	98			
188	3565.K03.G243	506865	59773	20.08	47.37	31.96	97			
241	3566.C15.G243	507073	60100		47.37	31.96	97			
259	3556.J14.G243	507064	60283		44.74	31.96	97			
269	3556.M02.G243	506876	62108		47.37	31.96	97			
275	3556.N05.G243	506940	58440		47.37	31.96	97			
334	3562.I02.G243	507338	58075		47.37	31.96	97			
466	3574.F10.G243	506318	58075		47.37	31.96	97			
472	3574.F10.G243	506318	58075		47.37	31.96	97			
472	3574.F10.G243	506318	58075	21.74	47.37	31.96	97			
621	3560.D18.G243	512389	60100		47.37	31.96	97			
621	3560.D18.G243	512389	60100		47.37	31.96	97			
636	3560.J01.G243	512107	60100		47.37	31.96	97			
644	3560.L10.G243	512253	60100		47.37	31.96	97			
689	3560.E08.G243	512688	60100		47.37	31.96	97			
671	3565.K14.G243	512700	60100		47.37	31.96	97			
681	3565.O10.G243	512640	60100		47.37	31.96	97			
688	3565.P07.G243	512553	60100		47.37	31.96	97			
708	3565.A23.G243	512553	60100		47.37	31.96	97			
708	3565.A23.G243	512553	60100		47.37	31.96	97			
713	3566.N04.G243	513228	60100		47.37	31.96	97			
713	3566.N04.G243	513228	60100		47.37	31.96	97			
718	3566.O06.G243	512960	60065		47.37	31.96	97			
721	3562.A09.G243	513378	60100		47.37	31.96	97			
740	3502.K05.G243	513340	60100		47.37	31.96	97			
750	3505.I18.G243	513390	60100		47.37	31.96	97			
798	3511.I04.G243	514468	60100		47.37	31.96	97			
798	3511.I04.G243	514468	60100		47.37	31.96	97			
820	3514.F11.G243	514951	59175	26.08	47.37	31.96	97			
820	3514.F11.G243	514951	59175	26.08	47.37	31.96	97			
822	3517.E18.G243	516411	59583		47.37	31.96	97			
822	3517.E18.G243	516411	59583		47.37	31.96	97			
825	3517.H18.G243	516417	59504		47.37	31.96	97			
825	3517.H18.G243	516417	59504		47.37	31.96	97			
835	3517.N14.G243	516361	57051		47.37	31.96	97			
835	3517.N14.G243	516361	57051		47.37	31.96	97			
835	3517.P11.G243	516345	56755		47.37	31.96	97			
838	3517.P12.G243	516361	60100		47.37	31.96	97			

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Table 16

SEQ ID	Seq Name	SpOBD	Breast Cancer Tumor/Normal >=2x Patients	Breast Cancer Tumor/Normal >=2x	Colon Cancer Tumor/Normal >=2x	Prostate Cancer Tumor/Normal >=2x	Prostate Cancer Tumor/Normal >=2x	Colon Unmatched Met >=2x Patients	Colon Matched Met/Normal >=2x Patients	Colon Matched Met/Normal >=2x	Colon Matched Met/Normal >=2x Patients	Colon Matched Met/Normal >=2x
836	3623.G02.G243.516810	57651										
837	3623.G02.G243.516811	57651										
846	3623.G02.G243.516828	27078			28.95	31.96	97	15.15	33			
865	3623.G01.G243.516858	33868	39.43	23		24.51	102	13.18	33			
872	3623.G01.G243.516859	35113	39.13	23		23.53	102	15.15	33			
876	3623.G01.G243.516861	59821	30.43	23		22.45	98					
876	3623.G01.G243.516861	59826	26.09	23	36.84	19				47.05	17	
887	3632.G01.G243.517319	26833				35.22	102	0.00	33			
889	3632.G02.G243.517627	60100				31.96	97					
901	3632.MT3.G243.517617	60100				31.96	97					
902	3632.MT9.G243.517613	57428				31.96	97					
903	3632.MT9.G243.517613	57428				31.96	97					
909	3632.A19.G243.517839	60100				31.96	97					
910	3632.A19.G243.517839	60100				31.96	97					
930	3633A.110.G243.518236	60100				31.96	97					
941	3643.G24.G243.516841	56804				25.77	97					
984	3661.K22.G243.519171	59763				20.41	98					
1043	3683.K19.G243.520821	25693				36.22	102	0.00	33			
1046	3664.P12.G243.520714	57708				23.47	98					
1081	3668.L23.G243.521064	57797				21.45	98					
1091	3754.S05.G243.532460	60100				28.47	102	0.00	33			
1101	3754.S05.G243.532460	60100				31.96	97					
1134	3755.C16.G243.533463	60100				31.96	97					
1139	3755.E12.G243.533461	57581				21.43	98					
1148	3765.G14.G243.533436	60100				31.96	97					
1172	3769.K05.G243.533678	58604				25.77	97					
1182	3762.A20.G243.534283	60233			47.37	18				41.16	17	
1216	3762.L18.G243.534272	60100				31.96	97					
1221	Ch02893.con.1	60100				31.96	97					
1238	Ch03948.con.1	60100				31.96	97					
1276	Ch059814.con.1	2451										
1296	Ch059814.con.1	2451										
1304	Ch0133440.con.1	24611										
1302	Ch0778070.con.1	60100										
1306	Ch0654673.con.1	57420				31.96	97					
1326	Ch0654673.con.1	60100				34.69	98					
1332	Ch01060739.con.1	58076				31.96	97					
1332	Ch01064483.con.1	60233			47.37	10						
1338	Ch01065336.con.1	25693				21.43	98					
1359	Ch01068930.con.1	27078				37.11	102	0.00	33			
1359	Ch01068930.con.1	27078			28.95	76		15.15	33			

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Table 16

SEQ ID	Seq Name	SpotID	Breast Cancer Tumor/Normal all >=2x Patients	Breast Cancer >=2x Patients	Colon Cancer Tumor/Normal all >=2x Patients	Colon Cancer Tumor/Normal all >=2x Patients	Prostate Cancer Tumor/Normal all >=2x Patients	Prostate Cancer Tumor/Normal all >=2x Patients	Colon Unmatched Met/Normal >=2x Patients	Colon Unmatched Met >=2x Patients	Colon Matched Met/Normal >=2x Patients	Colon Matched Met >=2x Patients	Colon Match Met Patients
1389	Ch11224379.con.1	42108		4474	76	31.68	97	63.64	33	32.76	35		
1408	Ch11226277.con.1	60100				31.68	102	0.00	33				
1413	Ch11250068.con.2	25944				49.02	102	0.00	33				
1418	Ch11250068.con.2	25944				20.41	98						
1421	Ch11262436.con.1	59664				25.77	97						
1440	NTN_00759252.3.10	53100		25.33	75			0.06	33	28.57	35		
1451	NTN_00926513.1	1368		34.29	35			56.67	30	28.57	7		
1451	NTN_00926513.1	25973		50.00	76			57.58	33	55.66	36		
1452	NTN_00926513.1	26568		59.21	76			66.67	33	44.44	36		
1452	NTN_00926513.2	59753				20.41	98						
1467	NTN_011512551.3.3	30086		22.67	75			6.66	33	25.00	36		
1468	NTN_011512551.3.3	39324		11.63	75			12.12	33	33.33	36		
1482	NTN_011512551.3.3	6974		69.74	76			78.79	33	66.67	36		
1482	NTN_02594513.2.1	27078		28.94	76			15.15	33				

We Claim:

1. An isolated polynucleotide comprising a nucleotide sequence which hybridizes under stringent conditions to a sequence selected from the group consisting of SEQ ID NOS: 1-1485, or complement thereof
2. An isolated polynucleotide comprising at least 15 contiguous nucleotides of a nucleotide sequence having at least 90% sequence identity to a sequence selected from the group consisting of SEQ ID NOS:1-1485, or complement thereof.
3. An isolated polynucleotide comprising at least 15 contiguous nucleotides of a nucleotide sequence selected from the group consisting of SEQ ID NOS:1-1485, or complement thereof.
4. The isolated polynucleotide of any one of claims 1-3, wherein the polynucleotide comprises at least 100 contiguous nucleotides of the nucleotide sequence or complement thereof.
5. The isolated polynucleotide of any one of claims 1-4, wherein the polynucleotide comprises at least 200 contiguous nucleotides of the selected nucleotide sequence or complement thereof.
6. An isolated polynucleotide comprising a nucleotide sequence of at least 90% sequence identity to a sequence selected from the group consisting of: SEQ ID NOS:1-1485 or complement therefore.
7. The isolated polynucleotide of claim 6, wherein the polynucleotide comprises a nucleotide sequence of at least 95% sequence identity to the selected nucleotide sequence.
8. The isolated polynucleotide of claim 6, wherein the polynucleotide comprises a nucleotide sequence that is identical to the selected nucleotide sequence.
9. A polynucleotide comprising a nucleotide sequence of an insert contained in a clone deposited as NRRL Accession No. B-30523, B-30524, B-30525, B-30526, B-30527, B-30528, B-30529, or B-30581.
10. An isolated cDNA obtained by the process of amplification using a polynucleotide comprising at least 15 contiguous nucleotides of a nucleotide sequence selected from the group consisting of SEQ ID NOS:1-1485.

11. The isolated cDNA of claim 10, wherein the polynucleotide comprises at least 25 contiguous nucleotides of the selected nucleotide sequence.

12. The isolated cDNA of claim 10, wherein the polynucleotide comprises at least 100 contiguous nucleotides of the selected nucleotide sequence.

13. The isolated cDNA of claims 10, 11, or 12, wherein amplification is by polymerase chain reaction (PCR) amplification.

14. An isolated recombinant host cell containing the polynucleotide according to claims 1, 2, 3, 6, 9, or 10.

15. An isolated vector comprising the polynucleotide according to claims 1, 2, 3, 6, 9, or 10.

16. A method for producing a polypeptide, the method comprising the steps of:  
culturing a recombinant host cell containing the polynucleotide according to claims 1, 2, 3, 6, 9, or 10, said culturing being under conditions suitable for the expression of an encoded polypeptide;  
and  
recovering the polypeptide from the host cell culture.

17. An isolated polypeptide encoded by the polynucleotide according to claims 1, 2, 3, 6, 9, or 10.

18. An isolated polypeptide comprising an amino acid sequence selected from the group consisting of SEQ ID NOS:1486-1542.

19. An antibody that specifically binds the polypeptide of claim 17 or 18.

20. A library of polynucleotides, wherein at least one of the polynucleotides comprises the sequence information of the polynucleotide according to claims 1, 2, 3, 6, 9, or 10.

21. The library of claim 20, wherein the library is provided on a nucleic acid array.

22. The library of claim 20, wherein the library is provided in a computer-readable format.

23. A method for detecting a cancerous cell, said method comprising:



detecting a level of a product of a gene in a test sample obtained from a cell of a subject, wherein said gene is identified by a sequence having at least 80% sequence identity to a sequence selected from a group consisting of SEQ ID NOS:1-1485, or a fragment thereof; and, comparing the level of said product to a control level of said gene product,  
5 wherein the presence of a cancerous cell is indicated by detection of said level and comparison to a control level of said gene product.

24. The method of claim 23, wherein said gene product is nucleic acid.

10 25. The method of claim 23, wherein said detecting step uses a polymerase chain reaction.

26. The method of claim 23, wherein said detecting step uses hybridization.

15 27. The method of claim 23, wherein said sample is a sample of prostate, colon or breast tissue.

28. A method for inhibiting a cancerous phenotype of a cell, said method comprising: contacting a mammalian cell with an agent for inhibition of a product of a gene, wherein  
20 said gene is identified by a sequence having at least 80% sequence identity to a sequence selected from a group consisting of SEQ ID NOS:1-1485, or a fragment thereof.

29. The method of claim 28, wherein said cancerous phenotype is aberrant cellular proliferation relative to a normal cell.

25 30. A method of treating a subject with cancer, said method comprising: administering to a subject a pharmaceutically effective amount of an agent, wherein said agent modulates the activity of a product of a gene identified by a sequence having at least 80% sequence identity to a sequence selected from a group consisting of SEQ ID  
30 NOS:1-1485, or a fragment thereof.

31. A method for identifying an agent that modulates a biological activity of a gene product differentially expressed in a cancerous cell as compared to a normal cell, said method comprising:

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contacting a candidate agent with a product of a gene encoded by a gene defined by a sequence having at least 80% sequence identity to a sequence selected from a group consisting of SEQ ID NOS:1-1485, or a fragment thereof; and

- 5 detecting modulation of a biological activity of the gene product relative to a level of biological activity of the gene product in the absence of the candidate agent.